

STIC Search Report Biotech-Chem Library

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TO: Ralph J Gitomer Location: 3d65 / 3c18 Thursday, May 11, 2006

Art Unit: 1655

Phone: 571-272-0916

Serial Number: 10 / 730070

From: Jan Delaval

Location: Biotech-Chem Library

Remsen 1a51

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Search Notes	



SEARCH REQUEST FORM

Scientific and Technical Information Center

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PTO-1590 (8-01)

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gitomer - 10 / 730070
=> fil medline
FILE 'MEDLINE' ENTERED AT 15:01:50 ON 11 MAY 2006
 FILE LAST UPDATED: 10 MAY 2006 (20060510/UP). FILE COVERS 1950 TO DATE.
 On December 11, 2005, the 2006 MeSH terms were loaded.
 The MEDLINE reload for 2006 is now (26 Feb.) available. For details
 on the 2006 reload, enter HELP RLOAD at an arrow prompt (=>).
 See also:
    http://www.nlm.nih.gov/mesh/
    http://www.nlm.nih.gov/pubs/techbull/nd04/nd04 mesh.html
    http://www.nlm.nih.gov/pubs/techbull/nd05/nd05 med data changes.html
    http://www.nlm.nih.gov/pubs/techbull/nd05/nd05 2006 MeSH.html
 OLDMEDLINE is covered back to 1950.
 MEDLINE thesauri in the /CN, /CT, and /MN fields incorporate the
 MeSH 2006 vocabulary.
 This file contains CAS Registry Numbers for easy and accurate
 substance identification.
=> d all tot
L29 ANSWER 1 OF 13
                        MEDLINE on STN
     97452104
                  MEDLINE
AN
DN
     PubMed ID: 9306875
ΤI
     Synovial fluid concentrations of the C-propeptide of type II
     collagen correlate with body mass index in primary knee
     osteoarthritis.
ΑU
     Kobayashi T; Yoshihara Y; Samura A; Yamada H; Shinmei M; Roos H; Lohmander
     L S
CS
     Department of Orthopaedic Surgery, National Defence Medical College,
     Tokorozawa, Japan.
SO
     Annals of the rheumatic diseases, (1997 Aug) Vol. 56, No. 8, pp.
     500-3.
     Journal code: 0372355. ISSN: 0003-4967.
CY
     ENGLAND: United Kingdom
     Journal; Article; (JOURNAL ARTICLE)
DΤ
LA
     English
FS
     Priority Journals
     199710
EM
ED
     Entered STN: 24 Oct 1997
     Last Updated on STN: 24 Oct 1997
     Entered Medline: 16 Oct 1997
AΒ
     OBJECTIVE: To explore in a cross sectional study in patients with primary
     knee osteoarthritis (OA) the relations between body mass index (BMI),
     disease stage, and the concentrations of a putative joint fluid marker of
     type II collagen synthesis, procollagen II
     C-propeptide. PATIENTS AND METHODS: The study included 142 patients with
     knee OA (median age 68, median BMI 24.1). OA was staged radiologically.
     The concentrations in synovial fluid of procollagen II
     C-propeptide were measured by a sandwich enzyme immunoassay.
     RESULTS: Joint fluid concentrations of procollagen II
```

C-propeptide were independently related to both OA stage and BMI (r =

C-propeptide were increased in knees with OA (median 3.7 ng/ml), compared with published reference values for knees in healthy adult volunteers

(median 1.3 ng/ml). The concentrations of procollagen II

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0.343, p < 0.0001 and r, = 0.253, p = 0.002, respectively). CONCLUSIONS:
     Joint fluid concentrations of this putative marker of collagen
     II synthesis are high in early and mid-stage OA, but decrease in end stage
     disease. In addition and for the first time it was shown that the
     concentrations in synovial fluid of procollagen II C-propeptide
     increase with increasing BMI in primary knee OA. The increased joint
     fluid values of this marker in patients with primary knee OA and a high
     BMI, may reflect increased rates of collagen synthesis in their
     joint cartilage and could relate to the previously shown increased risk
     for disease progression in such patients.
     Check Tags: Female; Male
      Adult
      Aged
      Aged, 80 and over
      Analysis of Variance
      Biological Markers: AN, analysis
     *Body Mass Index
     *Calcium-Binding Proteins: AN, analysis
       *Collagen: AN, analysis
        Collagen Type II
      Cross-Sectional Studies
      Disease Progression
      Humans
      Immunoenzyme Techniques
     *Knee Joint
      Knee Joint: RA, radiography
     Middle Aged
     *Osteoarthritis: ME, metabolism
     Osteoarthritis: RA, radiography
     *Protein Precursors: AN, analysis
      Regression Analysis
      Research Support, Non-U.S. Gov't
     Risk Factors
     *Synovial Fluid: CH, chemistry
     9007-34-5 (Collagen)
     0 (Biological Markers); 0 (Calcium-Binding Proteins); 0 (Collagen
     Type II); 0 (Protein Precursors); 0 (chondrocalcin)
                        MEDLINE on STN
L29
    ANSWER 2 OF 13
     97343445
                  MEDLINE
     PubMed ID: 9200001
     Measurement of bone degradation products in serum using antibodies
     reactive with an isomerized form of an 8 amino acid sequence of the
     C-telopeptide of type I collagen.
     Bonde M; Garnero P; Fledelius C; Qvist P; Delmas P D;
     Christiansen C
     Osteometer Bio Tech A/S, Herlev, Denmark.
     Journal of bone and mineral research : the official journal of the
     American Society for Bone and Mineral Research, (1997 Jul) Vol.
     12, No. 7, pp. 1028-34.
     Journal code: 8610640. ISSN: 0884-0431.
     United States
     Journal; Article; (JOURNAL ARTICLE)
     English
     Priority Journals
     199709
     Entered STN: 22 Sep 1997
     Last Updated on STN: 22 Sep 1997
     Entered Medline: 9 Sep 1997
     An enzyme-linked immunosorbent assay for measuring type I collagen
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degradation products in serum (S-ELISA) was developed. The assay uses a high affinity polyclonal antibody which reacts with an isomerized form of an 8 amino acid sequence of the C-telopeptides of type I collagen (EKAHD-beta-GGR). Cross-reactivity to a nonisomerized synthetic peptide form of the 8 amino acid sequence is less than 0.2%. Values obtained in a group of premenopausal women (age, 33.3) +/- 3.11 years) were 69 +/- 24 ng/ml(n = 22). In a group of early postmenopausal women (age, 51.8 +/- 1.88 years) values obtained were 125 +/- 43 ng/ml (n = 46), which represents an increase of 81% (p < 0.001). Values found in untreated patients with Paget's disease were 234 +/- 95 ng/ml (n = 15), and for primary hyperparathyroidism we found 335 +/- 82 ng/ml (n = 10). Intravenous administration of a bisphosphonate (Pamidronate) to Paget's disease patients for 3 days was reflected in the S-ELISA by a decrease in the values of 55% when compared with values before treatment (n = 15). Following treatment with another bisphosphonate (Alendronate) for 6 months, values were decreased to 48 +/-19 ng/ml (n = 12), which corresponds to a 62% decrease. Clinical results presented in this context support that the assay is a sensitive and specific index of bone resorption. It may, therefore, prove useful in the follow up of treatment of patients with metabolic bone diseases and in the clinical investigation of osteoporosis. Check Tags: Female Adult Alendronate: TU, therapeutic use Amino Acid Sequence Antibodies Biological Markers: BL, blood *Bone Resorption: BL, blood *Collagen: BL, blood Collagen: CH, chemistry Collagen: IM, immunology Cross Reactions Diphosphonates: TU, therapeutic use *Enzyme-Linked Immunosorbent Assay: MT, methods Enzyme-Linked Immunosorbent Assay: SN, statistics & numerical data Humans Hyperparathyroidism: BL, blood Menopause: BL, blood Menstruation: BL, blood Middle Aged Molecular Sequence Data Osteitis Deformans: BL, blood Osteitis Deformans: DT, drug therapy Osteoporosis: BL, blood *Peptides: BL, blood Peptides: CH, chemistry Peptides: IM, immunology Sensitivity and Specificity 40391-99-9 (pamidronate); 66376-36-1 (Alendronate); 9007-34-5 (Collagen) 0 (Antibodies); 0 (Biological Markers); 0 (Diphosphonates); 0 (Peptides); 0 (collagen type I trimeric cross-linked peptide) ANSWER 3 OF 13 MEDLINE on STN 97272754 MEDLINE PubMed ID: 9127460 Development of a monoclonal antibody to urinary degradation products from the C-terminal telopeptide alpha 1 chain of type I collagen. Application in an enzyme immunoassay and comparison to CrossLaps

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ELISA.
     Fledelius C; Kolding I; Qvist P; Bonde M; Hassager C; Reginster
ΑU
     J Y; Hejgaard J; Frookiaer H; Christiansen C
CS
     Osteometer BioTech AS, Herlev, Denmark.
SO
     Scandinavian journal of clinical and laboratory investigation, (1997
     Feb) Vol. 57, No. 1, pp. 73-83.
     Journal code: 0404375. ISSN: 0036-5513.
CY
     Norway
DT
     (CLINICAL TRIAL)
     (CONTROLLED CLINICAL TRIAL)
     Journal; Article; (JOURNAL ARTICLE)
LA
     English
FS
     Priority Journals
EM
     199706
ED
     Entered STN: 16 Jul 1997
     Last Updated on STN: 3 Mar 2000
     Entered Medline: 30 Jun 1997
AΒ
     A monoclonal antibody MADA7 was raised against a synthetic peptide having
     a sequence (EKAHDGGR) specific for a part of the C-telopeptide alpha 1
     chain of type I collagen. MAbA7 was labelled with horseradish
     peroxide and used in a competitive one-step enzyme-linked immunosorbent
     assay (ELISA) for measurement of urinary type I collagen
     degradation products. The assay was technically evaluated and preliminary
     clinical data are presented. The measuring range was 200-7000 micrograms
     1-1 with a detection limit of 25 micrograms 1-1. Within-run and total CVs
     were 5.5 and 8.0%, respectively. Analytical recovery averaged 96.6% +/-
     5.3 (mean +/- 1SD). Values obtained in the ELISA were highly
     correlated (r = 0.93) to values obtained by a commercially available assay
     (CrossLaps ELISA) known to measure urinary degradation
     products derived from the C-telopeptide of type I collagen
     reflecting the rate of bone resorption. Investigation of the urinary
     fragments responsible for the immunological response in the two assays
     revealed, however, that they are not identical. Values obtained in urine
     samples from postmenopausal women (n = 108) and patients with Paget's
     disease (n = 6) increased 43\% (p < 0.01) and 28-fold (p < 0.001),
     respectively, when compared to a premenopausal level (n = 50). A decrease
     in the urinary concentrations of 67\% (p < 0.01) was seen after 6 months in
     urine samples from postmenopausal women (n = 13) receiving hormone
     replacement therapy (HRT) compared to a group receiving placebo (n = 9).
     Likewise, the urinary concentrations decreased 88% (p < 0.001) in early
     postmenopausal women receiving bisphosphonate therapy (n = 11) for a
     period of 9 months compared to a group receiving placebo (n = 12).
     results suggest that the monoclonal antibody and the new assay may be
     useful for further investigations of the physiological and clinical
     importance of type I collagen degradation.
CT
    Check Tags: Female
     Adult
     Aged
     Amino Acid Sequence
     Antibodies, Monoclonal: BI, biosynthesis
     *Antibodies, Monoclonal: CH, chemistry
      Antibodies, Monoclonal: IP, isolation & purification
      Antibody Specificity
      Bone Resorption: IM, immunology
      Bone Resorption: UR, urine
        Collagen: CH, chemistry
       *Collagen: IM, immunology
       *Collagen: UR, urine
      Comparative Study
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Enzyme-Linked Immunosorbent Assay

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Humans
      Immunoenzyme Techniques
      Middle Aged
      Peptide Fragments: CS, chemical synthesis
     *Peptide Fragments: IM, immunology
     *Peptide Fragments: UR, urine
      Peptides: CH, chemistry
     *Peptides: IM, immunology
     *Peptides: UR, urine
      Postmenopause
      Premenopause
RN
     9007-34-5 (Collagen)
CN
     0 (Antibodies, Monoclonal); 0 (Peptide Fragments); 0 (Peptides); 0 (
     collagen type I trimeric cross-linked peptide)
L29 ANSWER 4 OF 13
                        MEDLINE on STN
ΑN
     97248522
                 MEDLINE
     PubMed ID: 9092508
DΝ
TΙ
    Characterization of urinary degradation products derived from type I
     collagen. Identification of a beta-isomerized Asp-Gly sequence
    within the C-terminal telopeptide (alphal) region.
ΑU
    Fledelius C; Johnsen A H; Cloos P A; Bonde M; Qvist P
CS
    Osteometer BioTech A/S, Herlev Hovedgade 207, DK-2730 Herlev, Denmark.
SO
    The Journal of biological chemistry, (1997 Apr 11) Vol. 272, No.
    15, pp. 9755-63.
    Journal code: 2985121R. ISSN: 0021-9258.
CY
    United States
DT
    Journal; Article; (JOURNAL ARTICLE)
LA
    English
FS
    Priority Journals
ΕM
    199705
ED
    Entered STN: 23 May 1997
    Last Updated on STN: 23 May 1997
    Entered Medline: 15 May 1997
AB
    The heterogeneity of urinary degradation products of C-terminal
    telopeptides derived from the alphal chain of human type I
    collagen was investigated and characterized. The urinary
    fragments characterized in this study consisted of two cross-linked (X)
    amino acid sequences derived from the C-terminal telopeptide (alphal) of
    type I collagen. Fragments containing the sequence EXAHDGGR,
    with a DG site being either nonisomerized (Asp-Gly) or beta-isomerized
     (betaAsp-Gly), were identified. Pyridinoline was detected among the
    pyridinium cross-links, but there was a dominance of deoxypyridinoline and
    a cross-link containing pyridinoline having a molecular weight identical
    with that of galactosyl pyridinoline. A nonfluorescent cross-link was
    also found. The concentration of fragments derived from the C-terminal
    telopeptide region of type I collagen containing the sequence
    Asp-Gly (alphaCTX) and/or betaAsp-Gly (betaCTX) was measured by
    enzyme-linked immunosorbent assays in urine and in collagenase
    digests of trabecular and cortical bone of young and old origin.
    shown that the urinary ratio between such fragments, alphaCTX/betaCTX, was
    higher in children compared with adults and that the ratio decreased with
    increasing age of bone. The results indicated that the C-terminal
    telopeptide fragments derived from type I collagen excreted into
    urine originated mainly from bone. In conclusion, it is demonstrated for
    the first time that the C-terminal telopeptide alphal chain of type I
    collagen contains an Asp-Gly site prone to undergo
    beta-isomerization and that the degree of beta-isomerization of this
    linkage apparently increases with increasing age of bone. These findings
    indicate that the ratio alphaCTX/betaCTX might be clinically important in
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diagnosing metabolic bone diseases.
CT
      Adult
      Amino Acid Sequence
      Antibodies, Monoclonal
      Aspartic Acid
      Bone and Bones: CH, chemistry
      Child
      Chromatography, High Pressure Liquid
       *Collagen: AN, analysis
       *Collagen: UR, urine
        Enzyme-Linked Immunosorbent Assay
      Glycine
      Humans
      Isomerism
     Molecular Sequence Data
     *Peptides: AN, analysis
RN
     56-40-6 (Glycine); 56-84-8 (Aspartic Acid); 9007-34-5 (Collagen)
CN
     0 (Antibodies, Monoclonal); 0 (Peptides); 0 (collagen type I
     trimeric cross-linked peptide)
L29
    ANSWER 5 OF 13
                        MEDLINE on STN
ΑN
     97027883
                  MEDLINE
DN
     PubMed ID: 8873970
TΤ
     High bone turnover is associated with low bone mass in both pre- and
     postmenopausal women.
ΑU
     Ravn P; Fledelius C; Rosenquist C; Overgaard K; Christiansen C
CS
     Center for Clinical and Basic Research, Ballerup, Denmark.
SO
     Bone, (1996 Sep) Vol. 19, No. 3, pp. 291-8.
     Journal code: 8504048. ISSN: 8756-3282.
CY
     United States
DT
     Journal; Article; (JOURNAL ARTICLE)
LA
     English
FS
     Priority Journals
EM
     199701
ΕD
     Entered STN: 19 Feb 1997
     Last Updated on STN: 19 Feb 1997
     Entered Medline: 29 Jan 1997
AB
     In 979 healthy women, aged 30-75 years, bone mass was measured by DXA in
     the lumbar spine and proximal femur, and by SXA in the distal forearm.
     Bone turnover was assessed by urinary CrossLaps (
     CrossLaps ELISA), a new assay which measures type I
     collagen degradation products in urine and by osteocalcin
     (two-site N-Mid hOsteocalcin ELISA), a new assay which measures
     the N-terminal-mid fragment (1-43) as well as the intact (1-49)
     osteocalcin (OCN-Mid) in serum. For comparison data on urinary
     hydroxyproline (fU Hpr/Cr) and serum, total alkaline phosphatase were
     included (AP). In premenopausal women below 50 years of age, the
     concentrations of the biochemical markers were stable with age.
     menopause CrossLaps and OCN-Mid increased abruptly to a level
     60% and 35% above the premenopausal mean values (p < 0.001).
     Premenopausal women in the highest quartiles, stratified according to the
     concentration of CrossLaps and OCN-Mid corrected for height and
     weight, had 6%-11% lower bone mass in all regions (p < 0.01) as compared
     to women in the lowest quartiles. CrossLaps and OCN-Mid
     corrected for height and weight correlated with bone mass in the spine and
     proximal femur, r = -0.13 to r = -0.28, p < 0.05. In postmenopausal
     women, the difference in bone mass between the highest and lowest
     quartiles was 8%-14% (p < 0.001). CrossLaps and OCN-Mid
     correlated with bone mass measured in all regions, r = -0.14 to r = -0.32,
     p < 0.05. The correlation between bone mass and AP and Fu Hpr/Cr was
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lower; r = -0.06 to r = -0.20 for premenopausal women, NS to p < 0.01, and r = -0.01 to r = -0.23, NS to p < 0.001 for postmenopausal women. conclusion, the present data indicate that high bone turnover is associated with a significantly lower bone mass in not only postmenopausal, but interestingly also in premenopausal women. In consistence with previous results, we found that bone turnover increased perimenopausally and in the early menopause. Check Tags: Female Adult Aged Analysis of Variance Biological Markers: CH, chemistry *Bone Density: PH, physiology *Bone Development: PH, physiology *Bone Resorption: PP, physiopathology Collagen: ME, metabolism Comparative Study Cross-Sectional Studies Enzyme-Linked Immunosorbent Assay Evaluation Studies Humans Linear Models Middle Aged Peptide Fragments: ME, metabolism *Postmenopause: PH, physiology *Premenopause: PH, physiology Reference Values 9007-34-5 (Collagen) 0 (Biological Markers); 0 (Peptide Fragments) L29 ANSWER 6 OF 13 MEDLINE on STN 97007889 MEDLINE PubMed ID: 8855148 Coated-tube radioimmunoassay for C-telopeptides of type I collagen to assess bone resorption. Bonde M; Fledelius C; Qvist P; Christiansen C Osteometer BioTech A/S, Herlev, Denmark. Clinical chemistry, (1996 Oct) Vol. 42, No. 10, pp. 1639-44. Journal code: 9421549. ISSN: 0009-9147. United States (CLINICAL TRIAL) (CONTROLLED CLINICAL TRIAL) Journal; Article; (JOURNAL ARTICLE) English Priority Journals 199612 Entered STN: 28 Jan 1997 Last Updated on STN: 28 Jan 1997 Entered Medline: 5 Dec 1996 We present a coated-tube RIA that is useful for assessment of bone resorption. The assay uses a monoclonal antibody raised against a linear 8-amino-acid sequence (EKAHDGGR) derived from the C-telopeptides of type I collagen. Within-run and total CVs were 4.4% and 5.3-6.2%, respectively, at concentrations of 1-7 mg/L (n = 4-20). Analytical recovery was 98% +/- 8% and dilution 97% +/- 7%. Values obtained in a group of 36 premenopausal women were 227 +/- 89.6 mg/mol creatinine. In a group of 141 postmenopausal women, the values obtained were 429 +/- 225 mg/mol creatinine, a highly significant increase of 89% (P <0.001) over the premenopausal value. In a double-blind placebo-controlled clinical study of these postmenopausal women receiving five different doses of a

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bisphosphonate, a significant decrease of RIA-measured C-telopeptide
     values was seen in all bisphosphonate-treated groups, after just 3 months.
     Values in urine samples from postmenopausal women assayed with the RIA
     (gamma) and the CrossLaps(TM) ELISA (x) agreed well:
     slope = 0.98 (95% confidence interval, 0.94-1.01), intercept = 0.34
     (0.25-0.43) mg/L, and Sylx = 0.93 mg/L (n = 678). We conclude that this
     RIA represents a valuable tool for assessing bone resorption.
CT
     Check Tags: Female
      Adult
      Aged
      Amino Acid Sequence
      Animals
      Antibodies, Monoclonal: IM, immunology
      Bone Density
      Bone Resorption: PC, prevention & control
     *Bone Resorption: UR, urine
        Collagen: IM, immunology
       *Collagen: UR, urine
      Diphosphonates: TU, therapeutic use
      Double-Blind Method
        Enzyme-Linked Immunosorbent Assay
      Humans
      Mice
      Mice, Inbred BALB C
      Middle Aged
      Peptide Fragments: IM, immunology
      Peptides: IM, immunology
     *Peptides: UR, urine
      Postmenopause: UR, urine
      Premenopause: UR, urine
     *Radioimmunoassay: MT, methods
      Reference Values
RN
     9007-34-5 (Collagen)
CN
     0 (Antibodies, Monoclonal); 0 (Diphosphonates); 0 (Peptide Fragments); 0
     (Peptides); 0 (collagen type I trimeric cross-linked peptide)
L29
    ANSWER 7 OF 13
                        MEDLINE on STN
AN
     95189878
                  MEDLINE
DN
     PubMed ID: 7883844
     Applications of an enzyme immunoassay for a new marker of bone resorption
ΤI
     (CrossLaps): follow-up on hormone replacement therapy and
     osteoporosis risk assessment.
ΑU
     Bonde M; Qvist P; Fledelius C; Riis B J; Christiansen C
CS
     Osteometer A/S, Rodovre, Denmark.
     The Journal of clinical endocrinology and metabolism, (1995 Mar)
     Vol. 80, No. 3, pp. 864-8.
     Journal code: 0375362. ISSN: 0021-972X.
CY
     United States
     (CLINICAL TRIAL)
     (CONTROLLED CLINICAL TRIAL)
     Journal; Article; (JOURNAL ARTICLE)
LA
     English
     Abridged Index Medicus Journals; Priority Journals
FS
EM
     199504
     Entered STN: 25 Apr 1995
     Last Updated on STN: 29 Jan 1999
     Entered Medline: 11 Apr 1995
AΒ
     An enzyme-linked immunosorbent immunoassay (ELISA) for a new
     marker of bone resorption (CrossLaps) was evaluated. The
    ELISA procedure determines degradation products of type I
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collagen in urine. Values obtained in the ELISA and in pyridinoline by high pressure liquid chromatography were correlated after a correction for creatinine. A high correlation was found (r = 0.77; n =81). A group of postmenopausal women (n = 180) showed an increase of more than 70% compared to values in premenopausal women (n = 104). Hydroxyproline was increased by 23%, osteocalcin by 52%, pyridinoline by 31%, and deoxypyridinoline by 50%. A highly significant decrease (60.7%) in the CrossLaps values was seen after 12 months in samples from patients receiving hormone replacement therapy compared to a placebo group. The spontaneous bone loss in an untreated group of women was determined by repeated forearm bone mass measurement over 24 months. Baseline values obtained in the CrossLaps ELISA were correlated to the rate of loss, yielding a highly significant r value of -0.61, indicating that CrossLaps might be a useful parameter for assessment of the risk of osteoporosis in postmenopausal women. Check Tags: Female Amino Acid Sequence Amino Acids: UR, urine Biological Markers *Bone Resorption: DI, diagnosis *Collagen: ME, metabolism Enzyme-Linked Immunosorbent Assay *Estrogen Replacement Therapy Follow-Up Studies Humans Middle Aged Molecular Sequence Data *Osteoporosis: ET, etiology 63800-01-1 (pyridinoline); 90032-33-0 (deoxypyridinoline); 9007-34-5 (Collagen) 0 (Amino Acids); 0 (Biological Markers) ANSWER 8 OF 13 MEDLINE on STN 95043378 MEDLINE PubMed ID: 7955372 Immunoassay for quantifying type I collagen degradation products in urine evaluated. Bonde M; Qvist P; Fledelius C; Riis B J; Christiansen C Center for Clinical and Basic Research, Ballerup, Denmark. Clinical chemistry, (1994 Nov) Vol. 40, No. 11 Pt 1, pp. 2022-5. Journal code: 9421549. ISSN: 0009-9147. Comment in: Clin Chem. 1994 Nov; 40(11 Pt 1):1994-5. PubMed ID: 7955367 United States Journal; Article; (JOURNAL ARTICLE) English Priority Journals 199412 Entered STN: 10 Jan 1995 Last Updated on STN: 10 Dec 2002 Entered Medline: 8 Dec 1994 An enzyme-linked immunosorbent assay (ELISA) for measuring type I collagen degradation products in urine < 3 h was evaluated. The measuring range was 0.5-10.5~mg/L with a detection limit of 0.2~mg/L. Within-run and total CVs were 5.3% and 6.6%, respectively. Analytical recovery averaged 100%. The mean (+/- SD) concentrations in urine samples from a healthy premenopausal population (n = 102) were 250 + /- 110 mg/molcreatinine (Cr). A group of healthy postmenopausal women (n = 410) gave a mean value of 416 +/- 189 mg/mol Cr. Values obtained in the ELISA correlated well (r = 0.83) to HPLC values for the established bone

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EΜ

AB

resorption marker deoxypyridinoline (n = 214), slightly better than the correlation to hydroxyproline measurements (r = 0.78, n = 421). conclude that the assay described here presents a useful tool for further elucidating the importance of type I collagen degradation products in urine. Check Tags: Female CTAmino Acid Sequence Chromatography, High Pressure Liquid Collagen: CH, chemistry *Collagen: UR, urine Comparative Study Creatinine: UR, urine Drug Stability *Enzyme-Linked Immunosorbent Assay: MT, methods Enzyme-Linked Immunosorbent Assay: SN, statistics & numerical data Freezing Humans Molecular Sequence Data Peptide Fragments: CH, chemistry *Peptide Fragments: UR, urine Postmenopause: UR, urine Premenopause: UR, urine Reference Values RN 60-27-5 (Creatinine); 9007-34-5 (Collagen) CN 0 (Peptide Fragments) L29 ANSWER 9 OF 13 MEDLINE on STN 94125564 MEDLINE AN DN PubMed ID: 8295344 ТΙ Evaluation of type IV collagen in patients with various thyroid disease. AU Senda Y; Nishibu M; Kawai K; Mizukami Y; Hashimoto T CS Central Clinical Laboratory, Kanazawa University School of Medicine. SO Rinsho byori. The Japanese journal of clinical pathology, (1993 Dec) Vol. 41, No. 12, pp. 1338-42. Journal code: 2984781R. ISSN: 0047-1860. CY Japan DTJournal; Article; (JOURNAL ARTICLE) LA Japanese FS Priority Journals 199402 EMEntered STN: 14 Mar 1994 Last Updated on STN: 14 Mar 1994 Entered Medline: 25 Feb 1994 AΒ Serum level of type IV collagen was measured in 104 patients with various thyroid disease, and the relationship between its level and thyroid hormone level was examined. The type IV collagen was measured by the method of one step sandwich enzyme immunoassay (EIA) using two distinct monoclonal antibodies recognized triple-helical (TH) domain and 7-S domain, respectively. The serum level of type IV collagen was significantly high in the hyperthyroid patients compared with that in normal controls, and a significant positive correlation was found between its value and thyroid hormone levels (T3, T4, FT3, FT4). The elevated level of type IV collagen in hyperthyroid patients was decreased to normal level, when they became to euthyroid after antithyroid drug therapy for hyperthyroidism. The serum level of type IV collagen was in normal range in hypothyroid patients, but the value was increased to high normal range after T4-replacement therapy for hypothyroidism. This evidence indicates that

the serum level of type IV collagen is closely related to thyroid hormone level in patient with various thyroid disease. collagen concentration might be one of the useful variables for evaluating the thyroid function, although its mechanism is not elucidated. CTCheck Tags: Female Adult *Collagen: BL, blood English Abstract Humans Hyperthyroidism: BL, blood Hypothyroidism: BL, blood Immunoenzyme Techniques *Thyroid Diseases: BL, blood Thyroid Hormones: BL, blood RN 9007-34-5 (Collagen) 0 (Thyroid Hormones) CN L29 ANSWER 10 OF 13 MEDLINE on STN AN 93047324 MEDLINE DN PubMed ID: 1330375 ΤI Concentration of serum laminin and type IV collagen in liver diseases assayed by a sandwich enzyme-immunoassay using monoclonal antibodies. ΑU Yokoya Y; Iwata K; Muragaki Y; Shiota C; Morimoto Y; Inoue M; Itoh H; Nishioka S; Ooshima A CS Department of Pathology, Wakayama Medical College, Japan. SO Clinica chimica acta; international journal of clinical chemistry, (1992 Sep 15) Vol. 210, No. 1-2, pp. 109-18. Journal code: 1302422. ISSN: 0009-8981. CY Netherlands DT Journal; Article; (JOURNAL ARTICLE) LA English FS Priority Journals EM 199212 Entered STN: 22 Jan 1993 Last Updated on STN: 22 Jan 1993 Entered Medline: 15 Dec 1992 AΒ Serum laminin (P1 fragment) and type IV collagen levels were determined in patients with hepatic disorders. The method was based on a sandwich enzyme-immunoassay using two monoclonal antibodies that recognize different epitopes of either laminin or type IV collagen molecule. Laminin and type IV collagen levels in the serum of patients with chronic hepatic disorders were higher as compared with those in healthy control subjects, with the increment of serum type IV collagen being far greater than that of laminin. Since type IV collagen and laminin are major basement membrane components, it is suggested that the higher levels of these peptides may reflect a so-called capillarization of the perisinusoidal wall encountered in hepatic fibrogenesis. The assay system used in this experiment is simple and sensitive and can be applied to clinical evaluation of hepatic fibrosis. CTAdolescent Adult Aged Aged, 80 and over *Antibodies, Monoclonal Carcinoma, Hepatocellular: BL, blood Carcinoma, Hepatocellular: CO, complications Child *Collagen: BL, blood Hepatitis: BL, blood

```
Humans
     *Immunoenzyme Techniques
     *Laminin: BL, blood
      Liver Cirrhosis: BL, blood
      Liver Cirrhosis: CO, complications
     *Liver Diseases: BL, blood
      Liver Neoplasms: BL, blood
      Liver Neoplasms: CO, complications
      Middle Aged
RN
     9007-34-5 (Collagen)
CN
     0 (Antibodies, Monoclonal); 0 (Laminin)
L29 ANSWER 11 OF 13
                         MEDLINE on STN
ΑN
     92335802
                  MEDLINE
DN
     PubMed ID: 1631498
TI
     Significance of serum type-IV collagen levels in various liver
     diseases. Measurement with a one-step sandwich enzyme
     immunoassay using monoclonal antibodies with specificity for
     pepsin-solubilized type-IV collagen.
ΑU
     Ueno T; Inuzuka S; Torimura T; Oohira H; Ko H; Obata K; Sata M; Yoshida H;
     Tanikawa K
CS
     Second Dept. of Medicine, Kurume University School of Medicine, Fukuoka,
SO
     Scandinavian journal of gastroenterology, (1992 Jun) Vol. 27,
     No. 6, pp. 513-20.
     Journal code: 0060105. ISSN: 0036-5521.
CY
     Norway
DΤ
     Journal; Article; (JOURNAL ARTICLE)
LA
     English
FS
     Priority Journals
EM
     199208
     Entered STN: 4 Sep 1992
ED
     Last Updated on STN: 4 Sep 1992
     Entered Medline: 14 Aug 1992
ΑB
     Serum type-IV collagen levels determined with a one-step
     sandwich enzyme immunoassay (EIA) using monoclonal antibodies with
     specificity for pepsin-solubilized type-IV collagen were
     compared with histologic changes in liver biopsy specimens from 107
     patients with various liver diseases. Serum type-IV collagen
     levels were increased in the groups with liver diseases compared with
     controls. The serum type-IV collagen levels in the group with
     alcoholic cirrhosis showed significantly higher values than the other
     groups (P less than 0.05). A significant positive correlation was found
     between the serum type-IV collagen level and the degree of
     fibrosis or cell infiltration in 107 patients. Immunolocalization of
     type-IV collagen was observed around blood vessels and bile
     ducts increased in number in the portal tracts, with cell infiltration and
     fibrosis, increased around vessels in fibrous septa, and sinusoidal walls
     of areas with cell infiltration or necrosis in hepatic lobules, and along
     the boundary between fibrous septa and hepatocytes. The present data
     indicate that serum type-IV collagen may be a sensitive marker
     for active fibrosis and that the elevation of serum type-IV
     collagen level primarily reflects the enhancement of type-IV
     collagen synthesis and deposition in the liver tissue at the stage
     of active fibrosis in liver disease.
     Check Tags: Female; Male
CT
      Adolescent
      Adult
      Aged
      Antibodies, Monoclonal
```

```
Collagen: AN, analysis
       *Collagen: BL, blood
      Hepatitis: BL, blood
      Hepatitis: PA, pathology
      Humans
      Immunoenzyme Techniques
      Immunohistochemistry
      Liver: CH, chemistry
      Liver: PA, pathology
     *Liver Diseases: BL, blood
      Liver Diseases: PA, pathology
      Liver Diseases, Alcoholic: BL, blood
      Liver Diseases, Alcoholic: PA, pathology
      Middle Aged
      Research Support, Non-U.S. Gov't
RN
     9007-34-5 (Collagen)
CN
     0 (Antibodies, Monoclonal)
    ANSWER 12 OF 13
L29
                         MEDLINE on STN
ΑN
     91011314
                  MEDLINE
DN
     PubMed ID: 1976752
TТ
     The occurrence and clinical significance of antibodies to type II
     collagen in sera and synovial fluid of Chinese patients with
     rheumatoid arthritis.
ΑU
     Chang M L; Chou C T; Lee C F
CS
     Department of Internal Medicine, Tri-Service General Hospital, National
     Defense Medical Center, Taipei, R.O.C.
SO
     Journal of the Formosan Medical Association = Taiwan yi zhi, (1990
    Apr) Vol. 89, No. 4, pp. 326-30.
     Journal code: 9214933. ISSN: 0929-6646.
CY
     TAIWAN: Taiwan, Province of China
DT
     Journal; Article; (JOURNAL ARTICLE)
LA
     Chinese
FS
     Priority Journals
EΜ
     199011
     Entered STN: 17 Jan 1991
     Last Updated on STN: 6 Feb 1995
     Entered Medline: 19 Nov 1990
AB
    Antibodies to type II collagen (Col II) in sera and synovial
     fluid (SF) were measured with an enzyme linked immunosorbent assay (
    ELISA) using a solid phase sandwich method. The
     subjects included: 42 patients with rheumatoid arthritis (RA); 31 cases of
     osteoarthritis (OA); 10 cases of gouty arthritis; 4 cases of ankylosing
     spondylitis (AS); 5 cases of systemic lupus erythematosus (SLE); and 44
     normal controls. The antigens used to detect antibodies against Col II
     were in native and heat-treated denatured forms, both of which were
     purified from chicken sternal cartilage by limited enzyme digestion and
     differential precipitation with salt. The reactivity to native type II
     collagen was generally higher than the reaction to the denatured
     collagen. In sera, significant higher levels of Col II were
     detected in the different arthritis groups when compared with the normal
     control group, with the exception of AS. In SF, the Col II was
     significantly higher in RA than it was in OA (p less than 0.001), while no
     difference was present between gout and OA (p less than 0.05). When
     native Col II was simultaneously measured in sera and SF among arthritics,
     positive rates were both higher among RA (65% and 58%, respectively).
     Positive rates were only higher in sera among OA (59% in sera and 3% in
     SF) and were both lower among gouty arthritis. The above findings show
     that the measurement of Col II is more important in SF than in sera.
CT
     *Arthritis, Rheumatoid: IM, immunology
```

```
*Autoantibodies: BL, blood
      China
       *Collagen: IM, immunology
      English Abstract
      Humans
     *Synovial Fluid: IM, immunology
RN
     9007-34-5 (Collagen)
CN
     0 (Autoantibodies)
L29
    ANSWER 13 OF 13
                         MEDLINE on STN
AN
     89337162
                  MEDLINE
DN
     PubMed ID: 2547537
     One step sandwich enzyme immunoassay for human type IV
ΤI
     collagen using monoclonal antibodies.
ΑU
     Obata K; Iwata K; Ichida T; Inoue K; Matsumoto E; Muragaki Y; Ooshima A
CS
     Department of Biotechnology, Fuji Chemical Industries, Ltd., Toyama,
SO
     Clinica chimica acta; international journal of clinical chemistry,
     (1989 May 31) Vol. 181, No. 3, pp. 293-303.
     Journal code: 1302422. ISSN: 0009-8981.
CY
     Netherlands
DT
     Journal; Article; (JOURNAL ARTICLE)
LA
     English
FS
     Priority Journals
EM
     198909
ED
     Entered STN: 9 Mar 1990
     Last Updated on STN: 9 Mar 1990
     Entered Medline: 20 Sep 1989
AΒ
     Monoclonal antibodies were used in one step sandwich enzyme
     immunoassay (one step sandwich EIA) for human serum
     immunoreactive type IV collagen. The one step sandwich
     EIA using either polystyrene ball or microplate was characterized by
     carrying out two immunoreactions simultaneously, type IV collagen
     reacting with both a monoclonal antibody as a solid phase and a
     horseradish peroxidase-labeled monoclonal antibody (Fab') against human
     type IV collagen as a conjugate. Sensitivity of one step
     sandwich EIA system by using either polystyrene ball or microplate
     was 0.22 ng per tube or 0.04 ng per well for type IV collagen,
     and linearity was obtained between 0.22-40 ng/tube or 0.04-20 ng per well,
     respectively. Both methods gave reproducible quantitative analysis of
     immunoreactive type IV collagen levels in the sera of patients
     with hepatocellular carcinoma and patients with liver cirrhosis, which
     were apparently higher than the levels in the sera of healthy subjects.
     Protein immunoblotting shows that the immunoreactive type IV
     collagen trapped in our present one step sandwich EIA
     system was not the 7-S and NC1 domains of type IV collagen.
CT
     *Antibodies, Monoclonal: AN, analysis
      Carcinoma, Hepatocellular: BL, blood
       *Collagen: BL, blood
        Collagen: IM, immunology
      Cross Reactions
      Humans
      Immunoblotting
      Immunoenzyme Techniques
      Liver Cirrhosis: BL, blood
      Liver Neoplasms: BL, blood
      Pepsin A
      Polystyrenes
     9007-34-5 (Collagen)
RN
CN
     0 (Antibodies, Monoclonal); 0 (Polystyrenes); EC 3.4.23.1 (Pepsin A)
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                            10 MAY 2006
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MOST RECENT DERWENT UPDATE:
                                200630
                                              <200630/DW>
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http://scientific.thomson.com/support/patents/coverage/latestupdates/
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http://www.stn-international.de/stndatabases/details/ipc reform.html and
http://scientific.thomson.com/media/scpdf/ipcrdwpi.pdf <<<
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=> d all abeq tech abex tot
L51 ANSWER 1 OF 4 WPIX COPYRIGHT 2006 THE THOMSON CORP on STN
AN
     2002-692261 [75]
                       WPIX
DNN N2002-546095
                        DNC C2002-195740
     Anti-human IV type collagen enzyme linked immunological
     quantitative determining kit and preparing method.
DC
     B04 D16 S03
     JIANG, P; MO, W
ΙN
PΑ
     (BEIJ-N) BEIJING CHEM REAGENT INST
CYC
                     A 19980722 (200275)*
PΤ
     CN 1188235
                                                    G01N033-53
ADT CN 1188235 A CN 1997-122057 19971219
PRAI CN 1997-122057
                          19971219
TC
    ICM G01N033-53
AR
    CN
          1188235 A UPAB: 20021120
     NOVELTY - The present invention relates to an antihuman IV type
     collagen protease linked immunoquantitative
     assay kit and its preparation method. It consists of
     enzyme scale plate and testing reagent, and uses human placenta to
     extract IV collagen, and adopts cell engineering--hybridoma
     technology to prepare monoclonal antibody, and uses enzyme
     linked immunosorbent principle to coat the monoclonal
     antibody on enzyme scale plate.
          DETAILED DESCRIPTION - The present invention relates to an antihuman
     IV type collagen protease linked immunoquantitative
     assay kit and its preparation method. It consists of
     enzyme scale plate and testing reagent, and uses human placenta to
     extract IV collagen, and adopts cell engineering--hybridoma
     technology to prepare monoclonal antibody, and uses enzyme
     linked immunosorbent principle to coat the monoclonal
     antibody on enzyme scale plate. It adopts double antibody
     sandwich method to quantitatively determine IV type
     collagen content in human serum. The invention can be used for
     diagnosis and treatment of fibrosis of liver and judgement after disease,
     and possesses good sensitivity, specificity and reproducibility and
     accuracy, so that it can meet requirement for clinical examination.
     Dwg.0/0
```

```
CPI EPI
FS
FΑ
     AB
MC
     CPI: B04-B04D4; B04-G01; B04-G21; B04-L01; B04-N02; B11-C07A4; B12-K04A;
          B14-N12; D05-A01A4; D05-A01B; D05-H09; D05-H11A
     EPI: S03-E14H4
    ANSWER 2 OF 4 WPIX COPYRIGHT 2006 THE THOMSON CORP on STN
L51
     1998-447376 [38]
ΑN
                        WPIX
DNN N1998-348672
                        DNC C1998-135811
ΤI
     Immunoassay kit containing two antibodies recognising coupled epitope(s)
     on collagen fragments - and new antibodies, for diagnosing
     arthritis etc., also prognosis and screening for anti-arthritic agents or
     inhibitors of matrix metallo-protease.
DC
     B04 D16 S03
IN
     CROUCHER, L J; HOLLANDER, A P
PA
     (UYSH-N) UNIV SHEFFIELD
CYC
     82
ΡI
     WO 9835235
                     Al 19980813 (199838) * EN
                                                59
                                                      G01N033-577
        RW: AT BE CH DE DK EA ES FI FR GB GH GM GR IE IT KE LS LU MC MW NL OA
            PT SD SE SZ UG ZW
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            GH GM GW HU ID IL IS JP KE KG KP KR KZ LC LK LR LS LT LU LV MD MG
            MK MN MW MX NO NZ PL PT RO RU SD SE SG SI SK SL TJ TM TR TT UA UG
            US UZ VN YU ZW
                     A 19980826 (199902)
     AU 9859560
                                                      G01N033-577
     EP 960339
                     Al 19991201 (200001)
                                           ΕN
                                                      G01N033-577
         R: CH DE FR GB IT LI SE
                    W 20010807 (200150)
     JP 2001511253
                                                59
                                                      G01N033-577
    WO 9835235 A1 WO 1998-GB304 19980130; AU 9859560 A AU 1998-59560 19980130;
ADT
     EP 960339 A1 EP 1998-902752 19980130, WO 1998-GB304 19980130; JP
     2001511253 W JP 1998-533982 19980130, WO 1998-GB304 19980130
     AU 9859560 A Based on WO 9835235; EP 960339 A1 Based on WO 9835235; JP
     2001511253 W Based on WO 9835235
PRAI GB 1997-2252
                          19970206
IC
     ICM G01N033-577
         CO7KO14-78; C07K016-18; C07K016-46; C12N005-10; C12N005-20;
          C12P021-08; G01N033-15; G01N033-50; G01N033-68
          9835235 A UPAB: 19981028
AB
     WO
     Immunoassay kit comprises two antibodies (Ab1 and Ab2), mono- or
     poly-clonal, or their fragments, that bind to two C-IIfree coupled
     epitopes (C-IIfree indicates any type II collagen fragment that
     is released from degraded cartilage). Also new are: (1) any Ab that binds
     to a coupled epitope on C-IIfree; (2) fragments of Ab; (3) bifunctional
     heteroantibodies (hAb) that bind to two C-IIfree coupled epitopes; (4)
     therapeutic agents identified by screening with the new kit; (5) cells, or
     cell lines, that express Ab or hAb and their fragments, and (6) isolated
     C-IIfree having at least 2 epitopes for production of Ab.
          The kits are designed for sandwich immunoassays,
     specifically enzyme-linked immunosorbent
     assay (ELISA), and C-IIfree is systemic (present in
     urine, serum or synovial fluid). The coupled epitopes comprise, or are
     present in, the N-terminal region of the alpha 1 type II collagen
     chain and can bind both Ab without mutual steric interference. The
     epitopes are conformational and/or contiguous and are separated by at
     least 2, up to 20, amino acids. Particularly Ab1 is immobilised on a
     support and/or Ab2 is labelled, particularly with biotin (used with an
     avidin-enzyme conjugate), radioisotope or enzyme
     (alkaline phosphatase or peroxidase). Abl is directed against epitope AH8,
     e.g. AH8MAb or AH8L1, and Ab2 is directed against AH12 (e.g. AH12L3), or
     vice versa.
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USE - The kits are used for therapy, diagnosis (e.g. routine
     screening for arthritis and other cartilage diseases, also to diagnose
     growth disorders), prognosis (e.g. monitoring progression of rheumatoid
     arthritis and osteoarthritis, or monitoring treatment with growth hormone)
     and for drug screening (to identify, and assess efficacy of,
     anti-arthritic agents and matrix metalloprotease inhibitors). hAb are used
     in immunoprecipitation assays.
          ADVANTAGE - C-IIfree, derived from the N-terminus of the alpha 1
     chain, have increased resistance to proteolysis, so can accumulate in vivo
     to a concentration that allows accurate measurement by immunoassay. Since
     they contain two or more epitopes, sandwich assays, which are
    more sensitive than inhibition assays and not subject to interference from
    collagen-binding proteins, can be developed.
     Dwg.0/20
    CPI EPI
    AΒ
    CPI: B04-G01; B11-C07; B12-K04A; B14-C09B; B14-N01; D05-A01A4; D05-A01B;
          D05-H08; D05-H09; D05-H11; D05-H15
    EPI: S03-E14H4
    ANSWER 3 OF 4 WPIX COPYRIGHT 2006 THE THOMSON CORP on STN
    1998-348692 [30]
                        WPIX
DNN
    N1998-272087
                        DNC C1998-107896
    Measurement of type I collagen resorption - by a
     sandwich assay for collagen degradation products.
    B04 D16 S03
    QVIST, P; ROSENQUIST, C; CHRISTGAU, S
     (OSTE-N) OSTEOMETER BIOTECH AS; (QVIS-I) QVIST P; (ROSE-I) ROSENQUIST C;
     (CHRI-I) CHRISTGAU S
    80
    WO 9826286
                     A2 19980618 (199830) * EN
                                                39
                                                      G01N033-53
        RW: AT BE CH DE DK EA ES FI FR GB GH GR IE IT KE LS LU MC MW NL OA PT
            SD SE SZ UG ZW
         W: AL AM AT AU AZ BA BB BG BR BY CA CH CN CU CZ DE DK EE ES FI GB GE
            GH HU ID IL IS JP KE KG KP KR KZ LC LK LR LS LT LU LV MD MG MK MN
            MW MX NO NZ PL PT RO RU SD SE SG SI SK SL TJ TM TR TT UA UG US UZ
           VN YU ZW
    AU 9857542
                    A 19980703 (199847)
                                                      G01N033-53
    EP 944833
                     A2 19990929 (199945) EN
                                                      G01N033-53
         R: AT BE CH DE DK ES FI FR GB GR IE IT LI LU MC NL PT SE
    JP 2001506000
                    W 20010508 (200131)
                                                40
                                                      G01N033-53
    EP 944833
                     B1 20010627 (200137)
                                          ΕN
                                                      G01N033-53
         R: AT BE CH DE DK ES FI FR GB GR IE IT LI LU MC NL PT SE
    DE 69705423
                     E 20010802 (200151)
                                                      G01N033-53
    ES 2160985
                     T3 20011116 (200201)
                                                      G01N033-53
    US 2003148272
                    A1 20030807 (200358)
                                                      C12Q001-68
    US 6660481
                     B2 20031209 (200381)
                                                      G01N033-53
    US 2004224375
                     A1 20041111 (200475)
                                                      G01N033-53
    WO 9826286 A2 WO 1997-EP6803 19971205; AU 9857542 A AU 1998-57542
    19971205; EP 944833 A2 EP 1997-953745 19971205, WO 1997-EP6803 19971205;
    JP 2001506000 W WO 1997-EP6803 19971205, JP 1998-526186 19971205; EP
    944833 B1 EP 1997-953745 19971205, WO 1997-EP6803 19971205; DE 69705423 E
    DE 1997-605423 19971205, EP 1997-953745 19971205, WO 1997-EP6803 19971205;
    ES 2160985 T3 EP 1997-953745 19971205; US 2003148272 A1 WO 1997-EP6803
    19971205, US 1999-319539 19990608; US 6660481 B2 WO 1997-EP6803 19971205,
    US 1999-319539 19990608; US 2004224375 A1 CIP of WO 1997-EP6803 19971205,
    CIP of US 1999-319539 19990608, US 2003-730070 20031209
FDT AU 9857542 A Based on WO 9826286; EP 944833 A2 Based on WO 9826286; JP
    2001506000 W Based on WO 9826286; EP 944833 B1 Based on WO 9826286; DE
    69705423 E Based on EP 944833, Based on WO 9826286; ES 2160985 T3 Based on
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FS

FA

MC

L51

AN

ΤI

DC

ΙN

PA

CYC

ADT

PΤ

EP 944833; US 6660481 B2 Based on WO 9826286; US 2004224375 A1 CIP of US 6660481

PRAI GB 1997-5687 19970319; GB 1996-25559 19961209

IC ICM C12Q001-68; G01N033-53

ICS C12N005-06; C12N005-16; G01N033-537; G01N033-543; G01N033-68

ICA C07K016-18

AB WO 9826286 A UPAB: 19980730

(A) A method is claimed for the measurement of the rate of type 1 collagen resorption comprising measuring in a sample the amount of a population of collagen fragments by a sandwich assay using a first antibody reactive with a first epitope located in the collagen amino acid sequence EKAHDGGR or in isomerised and/or racemised variants and a second antibody reactive with a second collagen epitope located in the fragments.

Also claimed are:

- (B) a method of conducting a sandwich assay comprising:
- (a) mixing a target antigen (TA) containing at least 2 antigenically similar epitopes with a first antibody reactive with both the epitopes, the first antibody is coupled to a capture moiety, and with a second antibody reactive with both the epitopes, the second antibody is coupled to a label, so as to form a first antibody TA second antibody sandwich; (b) capturing the sandwich to a capture substrate having an affinity for the capture moiety of the first antigen, and
- (c) detecting the capture of the **sandwich** by detection of the label of the second antibody;
- (C) a **sandwich** assay for **collagen** degradation products in which antibodies of identical specificity are used on both sides of the **sandwich**, and
- (D) a method of measurement of the concentration of **collagen** degradation products in a sample comprising conducting a **sandwich** assay using first and second immunological binding partners (which may be the same or different) each being immunologically reactive and an epitope in an N-terminal telopeptide fragment produced upon **collagen** degradation in vivo.
- USE The methods can be used to determine the metabolic status of tissues which generate **collagen**-derived peptides and isomerised and/or racemised peptide analogues when degradation occurs. They can be used to assess an abnormal condition of a subject by indicating excessive bone resorption. This may show the presence of an osteoporotic condition or the metastatic progress of a malignancy. Other conditions characterised by excessive bone resorption include Paget's disease and hyperparathyroidism.

Dwg.0/3

FS CPI EPI

FA AB

MC CPI: B04-B04C; B04-G01; B04-N02; B11-C07; B12-K04A; D05-H09; D05-H11A EPI: S03-E14H4

L51 ANSWER 4 OF 4 WPIX COPYRIGHT 2006 THE THOMSON CORP on STN

AN 1989-263792 [36] WPIX

DNN N1989-201270 DNC C1989-117142

TI Human type IV collagen determn. by sandwich enzyme immunoassay - for diagnosis of liver diseases such as hepatoma or chronic hepatitis.

DC B04 D16 J04 S03

IN INOUE, K; IWATA, K; OBATA, K; OSHIMA, A

PA (FUJY) FUJI YAKUHIN KOGYO KK; (FUJY) FUJI PHARM IND CO LTD

CYC 6

PI WO 8907761 A 19890824 (198936) * JA 59

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RW: DE FR GB IT
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     JP 02001553
                     A 19900105 (199007)
     EP 401370
                     A 19901212 (199050)
         R: DE FR GB IT
     JP 06077017
                    B2 19940928 (199437)
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                    A 19950317 (199520)
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                     E 19950629 (199531)
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    WO 8907761 A WO 1989-JP161 19890217; JP 02001553 A JP 1989-36111 19890217;
     EP 401370 A EP 1989-902540 19890217; JP 06077017 B2 JP 1989-36111
     19890217; EP 401370 A4 EP 1989-902540
                                                   ; JP 07072148 A Div ex JP
     1989-36111 19890217, JP 1993-252053 19890217; EP 401370 B1 EP 1989-902540
     19890217, WO 1989-JP161 19890217; DE 68922846 E DE 1989-622846 19890217,
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     JP 06077017 B2 Based on JP 02001553; EP 401370 B1 Based on WO 8907761; DE
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                          19880219; JP 1989-36111
     5.Jnl.Ref; DE 3115115; FR 2481318; GB 2074727; JP 57016355; JP 63246396;
     8.Jnl.Ref
     A61K039-39; C12P021-08; G01N033-53
TC
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          8907761 A UPAB: 19930923
     Assay of the central triple helix moiety of human type IV collagen
     is carried out by a single-stage sandwich enzyme immunoassay
     using a monoclonal antibody for a specific part of pepsin-modified human
     type IV collagen (pref. the collagen 7-S domain). The
     method uses as reagent solution an enzyme-labelled antibody recognising the
     central triple helix moiety of human type IV collagen; and as
     solid carrier to which is bound a monoclonal antibody recognising the
     central triple helix moiety of human type collagen. The reagent
     solution is added to the sample and after reaction the carrier-bound antibody
     is added and the enzyme activity measured.
          USE/ADVANTAGE - Simple and accurate diagnosis of liver disorders such
     as hepatitis, hepatoma and liver calcification.
     2/11
     CPI EPI
FS
FA
     AB; GI; DCN
     CPI: B04-B02C2; B04-B04A6; B04-B04C5; B04-B04D4; B05-C08; B10-B01A;
          B11-C07A4; B11-C07A6; B12-K04A; D05-A02A; D05-H09; D05-H11; J04-B01
     EPI: S03-E14H4
           401370 B UPAB: 19950630
ABEQ EP
     A method for the quantitation of the major central triple-helical region
     of human tye IV collagen by way of an ezyme immunoassay based on
     the sandwich technique using monoclonal antibodies against
     pepsin-solubilized human type IV collagen, in which (a) a sample
     solution prepared by diluting a sample to be assayed with a solution in a
     buffer of an enzyme-labelled antibody obtained by labelling with an enzyme
     a monoclonal antibody which reacts with the major central triple-helical
     region of pepsin-solubilised human type IV collagen and (b) an
     antibody-coated solid phase comprising a monoclonal antibody, bound to a
     solid phase carrier, which reacts only with pepsin-solubilised human type
     IV collagen are used, and which comprises mixing the
     antibody-coated solid phase (b) with the solution (a) to cause an
     immunoreaction to occur among the human type IV collagen present
     in the aid sample to be assayed, the said enzyme-labelled antibody and the
     antibody bound to the said solid phase carrier, isolating the solid phase
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carrier and measuring the enzyme activity bound to the solid phase thereby to quantitate the major central triple-helical region of human type IV ${f collagen}$. Dwg.0/11

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This file contains CAS Registry Numbers for easy and accurate substance identification.

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L81 ANSWER 1 OF 29 HCAPLUS COPYRIGHT 2006 ACS on STN
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AN 2004:964715 HCAPLUS

DN 141:408325

ED Entered STN: 12 Nov 2004

TI Sandwich assays for collagen fragments

IN Rosenquist, Christian; Qvist, Per; Christgau, Stephan

PA Den.

SO U.S. Pat. Appl. Publ., 14 pp., Cont.-in-part of U.S. Ser. No. 319,539. CODEN: USXXCO

DT Patent

LA English

IC ICM G01N0033-53 ICS G01N0033-53

G01N0033-537; G01N0033-543

INCL 435007930

CC 9-2 (Biochemical Methods)

Section cross-reference(s): 14

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                        G01N033/53; G01N033/543; G01N033/68R
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                 NCL
                        435/006.000
                 ECLA
                        G01N033/53; G01N033/543; G01N033/68R
AB
     Type II collagen degradation is measurable using
     a sandwich immunoassay in which a single antibody specific for
     the amino acid sequence EKGPDP is used to form each side of antibody-
     collagen fragment-antibody sandwich complexes and the
     amount of said complexes is measured.
ST
     sandwich assay collagen fragment
IT
     Disease, animal
        (arthropathy; sandwich assays for collagen
        fragments)
IT
     Joint, anatomical
        (disease; sandwich assays for collagen fragments)
IT
     Immunoassay
        (enzyme-linked immunosorbent
        assay; sandwich assays for collagen
        fragments)
IT
     Arthritis
     Blood serum
     Human
     Immunoassay
     Urine analysis
        (sandwich assays for collagen fragments)
     Collagens, analysis
     RL: ANT (Analyte); BSU (Biological study, unclassified); ANST (Analytical
     study); BIOL (Biological study)
        (sandwich assays for collagen fragments)
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ΙT
     Antibodies and Immunoglobulins
      RL: BSU (Biological study, unclassified); BIOL (Biological study)
         (sandwich assays for collagen fragments)
IT
      252578-68-0
      RL: ANT (Analyte); BSU (Biological study, unclassified); ANST (Analytical
      study); BIOL (Biological study)
         (collagen type II amino acid fragment;
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IT
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      154093-32-0
                     154093-34-2
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      791064-81-8 791313-09-2
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         (unclaimed sequence; sandwich assays for collagen
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L81 ANSWER 2 OF 29 HCAPLUS COPYRIGHT 2006 ACS on STN
AN
     2000:639145 HCAPLUS
DN
     133:234747
ED
    Entered STN: 14 Sep 2000
TΙ
    Assaying protein fragments in body fluids
IN
     Qvist, Per; Bonde, Martin
PA
     Osteometer Biotech A/s, Den.
SO
     U.S., 19 pp., Cont.-in-part of U.S. Ser. No. 913,806.
     CODEN: USXXAM
DT
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     English
IC
     ICM G01N0033-53
INCL 435007930
CC
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US 6323314 B1 20011127 US 2000-500811 20000210 <--
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US 6420125 B1 20020716 US 2000-641756 20000821 <--
US 2003119058 A1 20030626 US 2002-58124 20020129 <--
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                         530/388.100
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                        C07K016/18; G01N033/68R
AΒ
     Type I collagen degradation products are assayed in a body fluid by
     conducting a competition immunoassay in which sample mols. compete with a
     peptide or isomerized peptide in binding to an immunol. binding partner
     for the peptide or isomerized peptide resp., wherein the peptide or
     isomerized peptide comprises the amino acids AHDGGR optionally extended at
     the N-terminal end with one or more amino acids that do not form a
     contiguous sequence with AHDGGR in type 1 collagen, and wherein
     D represents aspartic acid or \beta-aspartic acid. The peptide C\left(X\right)n
     AHDGGR, where X is any amino acid and n is preferably from 4 to 6 is
     provided for use in such assays and in diagnostic assay kits.
ST
     assaying protein fragment body fluid
ΙT
     Immunoassay
        (Competition; assaying protein fragments in body fluids)
ΙT
     Antiserums
     Body fluid
     Diagnosis
```

```
Test kits
        (assaying protein fragments in body fluids)
    Amino acids, analysis
TT
    Peptides, analysis
    RL: ARU (Analytical role, unclassified); ANST (Analytical study)
        (assaying protein fragments in body fluids)
IT
     Proteins, general, analysis
    RL: ANT (Analyte); ANST (Analytical study)
        (fragments; assaying protein fragments in body fluids)
    Antibodies
IT
    RL: ARG (Analytical reagent use); ANST (Analytical study); USES (Uses)
        (monoclonal; assaying protein fragments in body fluids)
IT
    Collagens, analysis
    RL: ANT (Analyte); ANST (Analytical study)
        (type I, degradation products.; assaying protein fragments in body fluids)
IΤ
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                  162929-64-8
                                187269-53-0 284682-09-3
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    292139-65-2
                  292139-66-3
    RL: ARU (Analytical role, unclassified); ANST (Analytical study)
        (assaying protein fragments in body fluids)
TΤ
    292840-87-0
                 292840-88-1 292840-89-2
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                                                              292840-92-7
    RL: PRP (Properties)
        (unclaimed protein sequence; assaying protein fragments in body fluids)
RE.CNT
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              THERE ARE 125 CITED REFERENCES AVAILABLE FOR THIS RECORD
RE
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L81 ANSWER 3 OF 29 HCAPLUS COPYRIGHT 2006 ACS on STN
AN
     1998:550563 HCAPLUS
     129:172764
DN
ED
     Entered STN: 31 Aug 1998
     Immunoassay for collagen type II fragments
TΙ
IN
     Hollander, Anthony Peter; Croucher, Lisa Jane
PΑ
     University of Sheffield, UK
SO
     PCT Int. Appl., 60 pp.
     CODEN: PIXXD2
DT
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LA
     English
TC
     ICM G01N0033-577
         G01N0033-68; C07K0016-18; C07K0016-46; C12P0021-08; C07K0014-78;
          C12N0005-20
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     Section cross-reference(s): 13, 14, 15
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     Provided are methods, kits and reagents for assaying for collagen
AB
     fragments, and to therapeutic, prognostic and diagnostic methods based
     thereon. Antibodies and monoclonal antibodies to \alpha-chain
     type II collagen peptides were prepared,
     characterized, and used in sandwich ELISAs.
ST
     immunoassay collagen type II fragment;
     monoclonal antibody type II collagen
     ELISA
IT
     Synovial fluid
        (anal. of; immunoassay for collagen type II
        fragments)
IT
     Antibodies
     RL: ARG (Analytical reagent use); BPR (Biological process); BSU
     (Biological study, unclassified); THU (Therapeutic use); ANST (Analytical
     study); BIOL (Biological study); PROC (Process); USES (Uses)
        (bifunctional hetero-, to collagen type II
        fragments; immunoassay for collagen type II
        fragments)
ΙT
    Antibodies
     RL: ARG (Analytical reagent use); BPR (Biological process); BSU
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     (Therapeutic use); ANST (Analytical study); BIOL (Biological study); PREP
     (Preparation); PROC (Process); USES (Uses)
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```
(biotinylated; immunoassay for collagen type
        II fragments)
ΙT
     Radioactive substances
        (conjugates with antibodies; immunoassay for collagen
        type II fragments)
IT
     Enzymes, biological studies
     RL: ARG (Analytical reagent use); BPR (Biological process); BSU
     (Biological study, unclassified); THU (Therapeutic use); ANST (Analytical
     study); BIOL (Biological study); PROC (Process); USES (Uses)
        (conjugates, with antibodies; immunoassay for collagen
        type II fragments)
TT
     Avidins
     RL: ARG (Analytical reagent use); BPR (Biological process); BSU
     (Biological study, unclassified); THU (Therapeutic use); ANST (Analytical
     study); BIOL (Biological study); PROC (Process); USES (Uses)
        (conjugates, with enzymes; immunoassay for collagen
        type II fragments)
ΙT
     Cartilage
     Cartilage
        (degeneration, type II fragments, antibodies to;
        immunoassay for collagen type II
        fragments)
ΤТ
     Immunoassay
        (enzyme-linked immunosorbent
        assay, sandwich; immunoassay for collagen
        type II fragments)
TΤ
     Blood analysis
     Diagnosis
     Drug screening
     Immunoassay
     Urine analysis
        (immunoassay for collagen type II
        fragments)
ΙT
     Immunoassay
        (immunoblotting; immunoassay for collagen type
        II fragments)
IT
     Immunoassay
        (immunohistochem., of osteoarthritic cartilage; immunoassay for
        collagen type II fragments)
IT
     Osteoarthritis
        (immunostaining of cartilage of; immunoassay for collagen
        type II fragments)
IT
     Cartilage
        (immunostaining of osteoarthritic; immunoassay for collagen
        type II fragments)
ΙT
     Antibodies
     RL: ARG (Analytical reagent use); BPN (Biosynthetic preparation); BPR
     (Biological process); BSU (Biological study, unclassified); THU
     (Therapeutic use); ANST (Analytical study); BIOL (Biological study); PREP
     (Preparation); PROC (Process); USES (Uses)
        (monoclonal, to collagen type II
        fragments; immunoassay for collagen type II
        fragments)
IT
     Protein sequences
        (of synthetic peptide standard; immunoassay for collagen
        type II fragments)
ΙT
     Epitopes
        (on collagen type II fragments,
        antibodies to; immunoassay for collagen type
        II fragments)
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IT
     Antiarthritics
        (screening for; immunoassay for collagen type
        II fragments)
ΙT
     Antibodies
     RL: ARG (Analytical reagent use); BPN (Biosynthetic preparation); BPR
     (Biological process); BSU (Biological study, unclassified); PUR
     (Purification or recovery); RCT (Reactant); THU (Therapeutic use); ANST
     (Analytical study); BIOL (Biological study); PREP (Preparation); PROC
     (Process); RACT (Reactant or reagent); USES (Uses)
        (to collagen type II fragments;
        immunoassay for collagen type II
        fragments)
     Collagens, analysis
IT
     RL: ANT (Analyte); BPR (Biological process); BSU (Biological study,
     unclassified); THU (Therapeutic use); ANST (Analytical study); BIOL
     (Biological study); PROC (Process); USES (Uses)
        (type II; immunoassay for collagen
        type II fragments)
TΤ
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     RL: ARU (Analytical role, unclassified); PRP (Properties); SPN (Synthetic
     preparation); ANST (Analytical study); PREP (Preparation)
        (amino acid sequence of, as synthetic peptide standard; immunoassay for
        collagen type II fragments)
TΨ
     72040-63-2
     RL: RCT (Reactant); RACT (Reactant or reagent)
        (antibody biotinylation with; immunoassay for collagen
        type II fragments)
TT
     58-85-5DP, Biotin, antibody conjugates
     RL: ARG (Analytical reagent use); BPR (Biological process); BSU
     (Biological study, unclassified); SPN (Synthetic preparation); THU
     (Therapeutic use); ANST (Analytical study); BIOL (Biological study); PREP
     (Preparation); PROC (Process); USES (Uses)
        (immunoassay for collagen type II
        fragments)
IT
     9001-78-9D, antibody conjugates
                                       9003-99-0D, Peroxidase, antibody
     conjugates
     RL: ARG (Analytical reagent use); BPR (Biological process); BSU
     (Biological study, unclassified); THU (Therapeutic use); ANST (Analytical
     study); BIOL (Biological study); PROC (Process); USES (Uses)
        (immunoassay for collagen type II
        fragments)
IT
     141907-41-7, Matrix metalloproteinase
     RL: ADV (Adverse effect, including toxicity); BIOL (Biological study)
        (inhibitors, screening for; immunoassay for collagen
        type II fragments)
IT
     211370-80-8D, conjugates with keyhole limpet hemocyanin
                                                                211370-81-9D,
     conjugates with keyhole limpet hemocyanin 211370-82-0D, conjugates with
     keyhole limpet hemocyanin
     RL: BPR (Biological process); BSU (Biological study, unclassified); PRP
     (Properties); BIOL (Biological study); PROC (Process)
        (peptide of type II collagen
        α-chain; immunoassay for collagen type
        II fragments)
RE.CNT
              THERE ARE 3 CITED REFERENCES AVAILABLE FOR THIS RECORD
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L81 ANSWER 4 OF 29 HCAPLUS COPYRIGHT 2006 ACS on STN
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1998:406135 HCAPLUS
ΑN
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    129:78831
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    Sandwich immunoassays for collagen type I fragments
IN
    Rosenquist, Christian; Qvist, Per
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    Osteometer Biotech A/S, Den.; Rosenquist, Christian; Qvist, Per
SO
    PCT Int. Appl., 40 pp.
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    9-10 (Biochemical Methods)
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AΒ
     Sandwich assays for collagen degradation products are
     conducted using an antibody of the same specificity on both sides of the
     sandwich or using a first antibody reactive with an epitope
     contained in the sequence EKAHDGGR and a second antibody which may be the
     same or different. New collagen fragments were discovered in
     human serum. Two monoclonal antibodies (MAbs) were prepared by the
     hybridoma method that were specific for EKAH-\beta D-GGR. One of the MAbs
     was biotinylated and the other was coupled to horseradish peroxidase.
                                                                              The
     labeled MAbs were used in a sandwich assay to test urine samples
     from post-menopausal women taken before and after nine months of treatment
     with bisphosphonate.
     sandwich immunoassay collagen type I; monoclonal
ST
     antibody collagen peptide
     Blood serum
TΤ
     Urine
        (collagen fragments in human; sandwich immunoassays
        for collagen type I fragments)
IT
     Thyroglobulin
     RL: BPR (Biological process); BSU (Biological study, unclassified); BUU
     (Biological use, unclassified); SPN (Synthetic preparation); BIOL
     (Biological study); PREP (Preparation); PROC (Process); USES (Uses)
        (conjugates, with collagen fragment; sandwich
        immunoassays for collagen type I fragments)
IT
     Immunoassay
        (enzyme-linked immunosorbent
        assay, sandwich; sandwich immunoassays for
        collagen type I fragments)
IT
     Antigens
     RL: BOC (Biological occurrence); BSU (Biological study, unclassified);
     BIOL (Biological study); OCCU (Occurrence)
        (in human blood and urine reactive with monoclonal antibody to
        collagen epitope; sandwich immunoassays for
        collagen type I fragments)
```

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Antibodies
TT
     RL: ARG (Analytical reagent use); BPR (Biological process); BSU
     (Biological study, unclassified); SPN (Synthetic preparation); THU
     (Therapeutic use); ANST (Analytical study); BIOL (Biological study); PREP
     (Preparation); PROC (Process); USES (Uses)
        (monoclonal, biotinylated, to collagen fragment;
        sandwich immunoassays for collagen type I fragments)
TT
     Antibodies
     RL: ARG (Analytical reagent use); BPR (Biological process); BSU
     (Biological study, unclassified); SPN (Synthetic preparation); THU
     (Therapeutic use); ANST (Analytical study); BIOL (Biological study); PREP
     (Preparation); PROC (Process); USES (Uses)
        (monoclonal, labeled, to collagen fragment; sandwich
        immunoassays for collagen type I fragments)
     Antibodies
IT
     RL: ARG (Analytical reagent use); BPN (Biosynthetic preparation); BPR
     (Biological process); BSU (Biological study, unclassified); RCT
     (Reactant); THU (Therapeutic use); ANST (Analytical study); BIOL
     (Biological study); PREP (Preparation); PROC (Process); RACT (Reactant or
     reagent); USES (Uses)
        (monoclonal; sandwich immunoassays for collagen
        type I fragments)
IT
     Bone, disease
        (osteopenia, bone loss; sandwich immunoassays for
        collagen type I fragments)
TΤ
     Menopause
        (postmenopause; sandwich immunoassays for collagen
        type I fragments)
ΙT
     Immunoassay
        (radioimmunoassay, sandwich; sandwich immunoassays
        for collagen type I fragments)
TΨ
        (resorption; sandwich immunoassays for collagen
        type I fragments)
IT
     Hybridoma
     Urine analysis
        (sandwich immunoassays for collagen type I
        fragments)
IT
    Antibodies
     RL: ARG (Analytical reagent use); BPR (Biological process); BSU
     (Biological study, unclassified); THU (Therapeutic use); ANST (Analytical
     study); BIOL (Biological study); PROC (Process); USES (Uses)
        (sandwich immunoassays for collagen type I
        fragments)
IT
     Immunoassay
        (sandwich; sandwich immunoassays for
        collagen type I fragments)
ΙT
     Collagens, analysis
     RL: ANT (Analyte); THU (Therapeutic use); ANST (Analytical study); BIOL
     (Biological study); USES (Uses)
        (type I; sandwich immunoassays for collagen type I
        fragments)
ΙT
     162929-64-8
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     unclassified); THU (Therapeutic use); ANST (Analytical study); BIOL
     (Biological study); PROC (Process); USES (Uses)
        (collagen epitope; sandwich immunoassays for
        collagen type I fragments)
IT
     13598-36-2D, Phosphonic acid, alkylidenebis-derivs.
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
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(post-menopausal women treated with; sandwich immunoassays
        for collagen type I fragments)
     9003-99-ODP, Peroxidase, conjugates with monoclonal antibody to
IT
     collagen fragment
     RL: ARG (Analytical reagent use); BPR (Biological process); BSU
     (Biological study, unclassified); SPN (Synthetic preparation); THU
     (Therapeutic use); ANST (Analytical study); BIOL (Biological study); PREP
     (Preparation); PROC (Process); USES (Uses)
        (sandwich immunoassays for collagen type I
        fragments)
IT
     9013-20-1D, Streptavidin, immobilized
     RL: ARU (Analytical role, unclassified); THU (Therapeutic use); ANST
     (Analytical study); BIOL (Biological study); USES (Uses)
        (sandwich immunoassays for collagen type I
        fragments)
IT
     187269-53-0DP, conjugates with thyroglobulin
     RL: BPR (Biological process); BSU (Biological study, unclassified); BUU
     (Biological use, unclassified); SPN (Synthetic preparation); BIOL
     (Biological study); PREP (Preparation); PROC (Process); USES (Uses)
        (sandwich immunoassays for collagen type I
        fragments)
IT
     72040-63-2
     RL: RCT (Reactant); RACT (Reactant or reagent)
        (sandwich immunoassays for collagen type I
        fragments)
L81
    ANSWER 5 OF 29 HCAPLUS COPYRIGHT 2006 ACS on STN
AN
     1998:309527 HCAPLUS
DN
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ED
     Entered STN: 28 May 1998
ΤI
     Establishment of sandwich ELISA for determination of
     serum human type IV collagen and its preliminary application
ΑU
     Shen, Yi; Fan, Weike
CS
     Department of Pathophysiology, Chongqing University of Medical Sciences,
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    CODEN: MIZAED; ISSN: 1000-8861
PB
    Mianyixue Zazhi Bianjibu
DT
     Journal
LA
    Chinese
CC
     15-3 (Immunochemistry)
     Section cross-reference(s): 9, 14
AΒ
    Monoclonal antibody and polyclonal antibody to human Type IV
     collagen were produced by hybridoma technique and conventional
     immunization. A sandwich type of ELISA for serum
     human type IV collagen was established. The sensitivity of the
     immunoassay was 16 ng/mL. The CV value of intra-assays was 10.02%, of
     inter-assays was 9.14%. The accuracy was 92.68%. No cross reaction was
     found with Type I collagen. The concns. of serum Type IV
     collagen from 50 adult healthy people were 31.06 Φ+ 18.81 ng
    ml-1 with a range of 5 - 73 ng ml-1. Serum concentration of type IV
     collagen from patients with liver cirrhosis and cancer was higher
     than that from the healthy controls.
ST
     type IV collagen monoclonal polyclonal antibody; liver cancer
     cirrhosis antibody collagen IV
IT
     Immunoassay
        (enzyme-linked immunosorbent
        assay; preparation of antibodies for sandwich
        ELISA determination of serum human type IV collagen and for
        diagnosis of liver cancer and cirrhosis)
```

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IT
    Liver, disease
        (fibrosis; preparation of antibodies for sandwich ELISA
       determination of serum human type IV collagen and for diagnosis of
       liver cancer and cirrhosis)
IT
    Antibodies
    RL: BPN (Biosynthetic preparation); THU (Therapeutic use); BIOL
     (Biological study); PREP (Preparation); USES (Uses)
        (monoclonal; preparation of antibodies for sandwich ELISA
       determination of serum human type IV collagen and for diagnosis of
       liver cancer and cirrhosis)
IT
    Blood serum
    Cirrhosis
    Liver, neoplasm
        (preparation of antibodies for sandwich ELISA determination of
       serum human type IV collagen and for diagnosis of liver
       cancer and cirrhosis)
ΙT
    Antibodies
    RL: BPN (Biosynthetic preparation); THU (Therapeutic use); BIOL
     (Biological study); PREP (Preparation); USES (Uses)
        (preparation of antibodies for sandwich ELISA determination of
       serum human type IV collagen and for diagnosis of liver
       cancer and cirrhosis)
ΙT
    Collagens, biological studies
    RL: BSU (Biological study, unclassified); THU (Therapeutic use); BIOL
     (Biological study); USES (Uses)
        (type IV; preparation of antibodies for sandwich ELISA
       determination of serum human type IV collagen and for diagnosis of
       liver cancer and cirrhosis)
L81 ANSWER 6 OF 29 HCAPLUS COPYRIGHT 2006 ACS on STN
AN
    1998:186591 HCAPLUS
DN
    128:241535
ΕĎ
    Entered STN: 30 Mar 1998
    Assay for detecting collagen degradation
ΤI
IN
    Te Koppele, Johannes Maria; Beekman, Bob
PA
    Nederlandse Organisatie voor Toegepast-Natuurwetenschappelijk Onderzoek
    TNO, Neth.
    Eur. Pat. Appl., 31 pp.
SO
    CODEN: EPXXDW
DT
    Patent
LA
    English
IC
    ICM G01N0033-68
    9-10 (Biochemical Methods)
    Section cross-reference(s): 14
FAN.CNT 1
    PATENT NO.
                      KIND DATE APPLICATION NO. DATE
    EP 829724 A1 19980318 EP 1996-202596 19960917 <--
        R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
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    JP 10185915
                       A2
                                       JP 1997-268200
                              19980714
                                                              19970916 <--
                                       US 1997-931820
                       Α
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PRAI EP 1996-202596 A
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CLASS
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               ICM
EP 829724
                      G01N0033-68
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                      G01N033/68R
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JP 10185915
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 US 6010863
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                 IPCI
                 IPCR
                        G01N0033-68 [I,A]; G01N0033-68 [I,C]
                 NCL
                        435/007.100; 435/007.900; 435/007.920; 435/007.940;
                        435/975.000; 436/518.000; 436/531.000
                 ECLA
                        G01N033/68R
AB
     A sandwich-type immunoassay for the detection and /or
     quantitation of collagen degradation products in biol. samples such
     as blood, serum, plasma, sputum and cell cultures. The immunoassay uses a
     first antibody directed at an epitope present on a collagen mol.
     at a distance of up to 165 amino acids from a collagen
     telopeptide crosslink site, and a second antibody directed at another
     epitope of the crosslinked collagen mol.
ST
     collagen degrdn product sandwich immunoassay sequence
IT
     Antibodies
     RL: ARG (Analytical reagent use); BPN (Biosynthetic preparation); BPR
     (Biological process); BSU (Biological study, unclassified); THU
     (Therapeutic use); ANST (Analytical study); BIOL (Biological study); PREP
     (Preparation); PROC (Process); USES (Uses)
        (collagen-specific; immunoassay for detecting
        collagen degradation)
IT
     Bone
        (degradation products; immunoassay for detecting collagen
        degradation)
TΤ
     Collagens, biological studies
     RL: BSU (Biological study, unclassified); BIOL (Biological study)
        (degradation products; immunoassay for detecting collagen
        degradation)
IT
     Antibodies
     RL: BPN (Biosynthetic preparation); BPR (Biological process); BSU
     (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological
     study); PREP (Preparation); PROC (Process); USES (Uses)
        (immobilized; immunoassay for detecting collagen degradation)
TΨ
     Animal tissue culture
     Blood
     Blood analysis
     Blood serum
     Protein sequences
     Sputum
     Test kits
        (immunoassay for detecting collagen degradation)
ΙT
     Immunoassay
        (sandwich; immunoassay for detecting collagen
        degradation)
TΤ
     Collagens, analysis
     RL: ANT (Analyte); BPR (Biological process); BSU (Biological study,
     unclassified); ANST (Analytical study); BIOL (Biological study); PROC
     (Process)
        (type I; immunoassay for detecting collagen degradation)
TT
     Collagens, analysis
     RL: ANT (Analyte); BPR (Biological process); BSU (Biological study,
     unclassified); ANST (Analytical study); BIOL (Biological study); PROC
     (Process)
        (type II; immunoassay for detecting
        collagen degradation)
ΙT
     Collagens, analysis
     RL: ANT (Analyte); BPR (Biological process); BSU (Biological study,
     unclassified); ANST (Analytical study); BIOL (Biological study); PROC
     (Process)
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(type III; immunoassay for detecting collagen degradation)
IT
     Collagens, analysis
     RL: ANT (Analyte); BPR (Biological process); BSU (Biological study,
     unclassified); ANST (Analytical study); BIOL (Biological study); PROC
     (Process)
        (type IV; immunoassay for detecting collagen degradation)
IT
     Collagens, analysis
     RL: ANT (Analyte); BPR (Biological process); BSU (Biological study,
     unclassified); ANST (Analytical study); BIOL (Biological study); PROC
     (Process)
        (type IX; immunoassay for detecting collagen degradation)
TT
     Collagens, analysis
     RL: ANT (Analyte); BPR (Biological process); BSU (Biological study,
     unclassified); ANST (Analytical study); BIOL (Biological study); PROC
     (Process)
        (type V; immunoassay for detecting collagen degradation)
IT
     Collagens, analysis
     RL: ANT (Analyte); BPR (Biological process); BSU (Biological study,
     unclassified); ANST (Analytical study); BIOL (Biological study); PROC
     (Process)
        (type VI; immunoassay for detecting collagen degradation)
IT
     Collagens, analysis
     RL: ANT (Analyte); BPR (Biological process); BSU (Biological study,
     unclassified); ANST (Analytical study); BIOL (Biological study); PROC
     (Process)
        (type X; immunoassay for detecting collagen degradation)
TΤ
     Collagens, analysis
     RL: ANT (Analyte); BPR (Biological process); BSU (Biological study,
     unclassified); ANST (Analytical study); BIOL (Biological study); PROC
     (Process)
        (type XI; immunoassay for detecting collagen degradation)
RE.CNT
             THERE ARE 4 CITED REFERENCES AVAILABLE FOR THIS RECORD
RF.
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(2) Fuji Yakuhin Kogyo Kk; EP 0401370 A 1990 HCAPLUS
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(4) Osteometer A S; WO 9508115 A 1995 HCAPLUS
L81 ANSWER 7 OF 29 HCAPLUS COPYRIGHT 2006 ACS on STN
AN
    1998:151309 HCAPLUS
DN
     128:164717
     Entered STN: 13 Mar 1998
ED
     Assaying D-amino acids in body fluids
TΙ
ΙN
     Fledelius, Christian; Cloos, Paul; Qvist, Per
     Osteometer Biotech A/S, Den.; Fledelius, Christian; Cloos, Paul; Qvist,
PΑ
     Per
SO
     PCT Int. Appl., 42 pp.
     CODEN: PIXXD2
DT
     Patent
LA
     English
TC
     ICM G01N0033-68
     ICS C07K0016-18; C12N0005-12; G01N0033-577; C07K0014-78
CC
     9-10 (Biochemical Methods)
FAN.CNT 1
     PATENT NO.
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                                            APPLICATION NO.
                                DATE
                                                                  DATE
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     WO 9808098
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                         A2
                                                                  19970812 <--
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            DK, EE, ES, FI, GB, GE, GH, HU, IL, IS, JP, KE, KG, KP, KR, KZ,
            LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL,
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PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG, US,
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             GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA,
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     AU 9739435
                          Α1
                                19980306
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                                                                   19970812 <--
     EP 922228
                          A2
                                19990616
                                            EP 1997-936704
                                                                   19970812 <--
         R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
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     JP 2000516721
                          Т2
                                20001212
                                            JP 1998-510349
                                                                   19970812 <--
                          В1
     US 6300083
                                20011009
                                            US 2000-242721
                                                                   20000110 <--
PRAI GB 1996-17616
                         Α
                                19960822
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     WO 1997-EP4372
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CLASS
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                       PATENT FAMILY CLASSIFICATION CODES
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 WO 9808098
                 ICM
                        G01N0033-68
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                        C07K0016-18; C12N0005-12; G01N0033-577; C07K0014-78
                 IPCI
                        G01N0033-68 [ICM, 6]; C07K0016-18 [ICS, 6]; C12N0005-12
                        [ICS, 6]; G01N0033-577 [ICS, 6]; C07K0014-78 [ICS, 6]
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                        C07K0014-435 [I,C]; C07K0014-78 [I,A]; C07K0016-18
                        [I,A]; C07K0016-18 [I,C]; G01N0033-68 [I,A];
                        G01N0033-68 [I,C]
                 ECLA
                        C07K014/78; C07K016/18; G01N033/68R
 AU 9739435
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 JP 2000516721
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                        G01N0033-53 [ICM,7]; G01N0033-68 [ICS,7]
 US 6300083
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                        G01N0033-53 [ICM, 7]
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                        G01N0033-68 [I,C]
                 NCL
                        435/007.100; 435/007.920; 435/007.930; 435/007.940;
                        435/007.950; 435/331.000; 436/518.000; 436/532.000;
                        530/323.000; 530/326.000; 530/327.000; 530/328.000;
                        530/329.000; 530/356.000; 530/388.100; 530/389.100
                        C07K014/78; C07K016/18; G01N033/68R
                 ECLA
AΒ
     The rate of degradation in vivo of a body protein is determined by measuring
the
     amount of a D-amino acid containing fragment of the protein in a body fluid
     using an antibody capable of discriminating between the D-amino acid
     containing fragment and its L-amino acid containing analog.
     assaying amino acid body fluid
ST
IT
     Body fluid
     Immunoassay
     Protein degradation
     Urine analysis
        (assaying D-amino acids in body fluids)
IT
     Collagens, analysis
     Enzymes, analysis
     Peptides, analysis
     RL: ANT (Analyte); ANST (Analytical study)
        (assaying D-amino acids in body fluids)
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IT
     Antibodies
     RL: ARG (Analytical reagent use); ANST (Analytical study); USES (Uses)
        (assaying D-amino acids in body fluids)
     Amino acids, analysis
ΙT
     RL: ARU (Analytical role, unclassified); ANST (Analytical study)
        (assaying D-amino acids in body fluids)
IT
     Immunoassay
        (enzyme-linked immunosorbent
        assay; assaying D-amino acids in body fluids)
TΤ
     Washing
        (solution; assaying D-amino acids in body fluids)
TΤ
     Amino acids, analysis
     RL: ANT (Analyte); ANST (Analytical study)
        (D-; assaying D-amino acids in body fluids)
IT
     1783-96-6, D-Aspartic acid
     RL: ANT (Analyte); ANST (Analytical study)
        (assaying D-amino acids in body fluids)
IT
     56-40-6, Glycine, analysis
     RL: ARU (Analytical role, unclassified); ANST (Analytical study)
        (assaying D-amino acids in body fluids)
L81 ANSWER 8 OF 29 HCAPLUS COPYRIGHT 2006 ACS on STN
AN
     1997:464042 HCAPLUS
DN
     127:146712
ED
     Entered STN: 24 Jul 1997
TΤ
    Measurement of bone degradation products in serum using antibodies
     reactive with an isomerized form of an 8 amino acid sequence of the
     C-telopeptide of type I collagen
ΑU
     Bonde, Martin; Garnero, Patrick; Fledelius, Christian; Qvist, Per
     ; Delmas, Pierre D.; Christiansen, Claus
CS
     Osteometer BioTech A/S, Herlev, Den.
SO
     Journal of Bone and Mineral Research (1997), 12(7), 1028-1034
     CODEN: JBMREJ; ISSN: 0884-0431
PB
     Blackwell
DΤ
     Journal
LA
     English
CC
     9-10 (Biochemical Methods)
     Section cross-reference(s): 6, 13, 14
AB
    An ELISA for measuring type I collagen degradation
    products in serum (S-ELISA) was developed. The assay uses a
     high affinity polyclonal antibody which reacts with an isomerized form of
     an 8 amino acid sequence of the C-telopeptides of type I collagen
     (EKAHD-\beta-GGR). Cross-reactivity to a nonisomerized synthetic peptide
     form of the 8 amino acid sequence is less than 0.2%. Values obtained in a
     group of premenopausal women (age, 33.3±3.11 yr) were 69±24 ng/mL.
     In a group of early postmenopausal women (age, 51.8±1.88 yr) values
     obtained were 125±43 ng/mL, which represents an increase of 81%.
     Values found in untreated patients with Paget's disease were 234±95
     ng/mL, and for primary hyperparathyroidism we found 335±82 ng/mL.
     Intervenous administration of a bisphosphonate (Pamidronate) to Paget's
     disease patients for 3 days was reflected in the S-ELISA by a
     decrease in the values of 55% when compared with values before treatment.
     Following treatment with another bisphosphonate (Alendronate) for 6 mo,
     values were decreased to 48\pm19~\mathrm{ng/mL}, which corresponds to a 62%
     decrease. Clin. results presented in this context support that the assay
     is a sensitive and specific index of bone resorption. It may, therefore,
     prove useful in the follow up of treatment of patients with metabolic bone
     diseases and in the clin. investigation of osteoporosis.
ST
     bone degeneration collagen telopeptide ELISA
ΙT
     Peptides, analysis
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jan delaval - 11 may 2006

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RL: ARU (Analytical role, unclassified); BPR (Biological process); BSU
     (Biological study, unclassified); ANST (Analytical study); BIOL
     (Biological study); PROC (Process)
        (C-telopeptides; measurement of bone degradation products in serum using
        ELISA)
ΙT
     Bone, disease
        (Paget's; measurement of bone degradation products in serum using
        ELISA)
ΙT
        (degradation; measurement of bone degradation products in serum using
        ELISA)
ΙT
     Immunoassay
        (enzyme-linked immunosorbent
        assay; measurement of bone degradation products in serum using
        ELISA)
IT
     Blood analysis
        (measurement of bone degradation products in serum using ELISA)
IT
     Hyperparathyroidism
        (primary; measurement of bone degradation products in serum using
        ELISA)
ΙT
        (resorption; measurement of bone degradation products in serum using
        ELISA)
TT
     Collagens, biological studies
     RL: BPR (Biological process); BSU (Biological study, unclassified); BIOL
     (Biological study); PROC (Process)
        (type I; measurement of bone degradation products in serum using
        ELISA)
IT
     187269-53-0
     RL: ANT (Analyte); BSU (Biological study, unclassified); ANST (Analytical
     study); BIOL (Biological study)
        (measurement of bone degradation products in serum using ELISA)
IT
     40391-99-9
                  66376-36-1, Alendronate
     RL: BSU (Biological study, unclassified); THU (Therapeutic use); BIOL
     (Biological study); USES (Uses)
        (measurement of bone degradation products in serum using ELISA)
L81 ANSWER 9 OF 29 HCAPLUS COPYRIGHT 2006 ACS on STN
     1997:257774 HCAPLUS
AN
DN
     126:328573
ED
     Entered STN: 21 Apr 1997
ΤI
     Characterization of urinary degradation products derived from type I
     collagen
ΑU
     Fledelius, Christian; Johnsen, Anders H.; Cloos, Paul A. C.; Bonde,
     Martin; Qvist, Per
CS
     Osteometer BioTech A/S, Herlev, DK-2730, Den.
SO
     Journal of Biological Chemistry (1997), 272(15), 9755-9763
     CODEN: JBCHA3; ISSN: 0021-9258
PΒ
     American Society for Biochemistry and Molecular Biology
DT
     Journal
LA
     English
CC
     13-3 (Mammalian Biochemistry)
     Section cross-reference(s): 6
AB
     The heterogeneity of urinary degradation products of C-terminal telopeptides
     derived from the \alpha 1 chain of human type I collagen was
     investigated and characterized. The urinary fragments characterized in
     this study consisted of two cross-linked (X) amino acid sequences derived
     from the C-terminal telopeptide (\alpha1) of type I collagen.
     Fragments containing the sequence EXAH-DGGR, with a DG site being either
     nonisomerized (Asp-Gly) or \beta-isomerized (\betaAsp-Gly), were
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identified. Pyridinoline was detected among the pyridinium crosslinks, but there was a dominance of deoxypyridinoline and a cross-link containing pyridinoline having a mol. weight identical with that of galactosyl pyridinoline. A nonfluorescent cross-link was also found. The concentration

of

fragments derived from the C-terminal telopeptide region of type I collagen containing the sequence Asp-Gly (αCTX) and/or $\beta Asp-Gly$ (βCTX) was measured by enzyme-linked immunosorbent assays in urine and in collagenase digests of trabecular and cortical bone of young and old origin. It was shown that the urinary ratio between such fragments, $\alpha CTX/\beta CTX$, was higher in children compared with adults and that the ratio decreased with increasing age of bone. The results indicated that the C-terminal telopeptide fragments derived from type I collagen excreted into urine originated mainly from bone. In conclusion, it is demonstrated for the first time that the C-terminal telopeptide $\alpha 1$ chain of type I collagen contains an Asp-Gly site prone to undergo β -isomerization and that the degree of β -isomerization of this linkage apparently increases with increasing age of bone. These findings indicate that the ratio $\alpha CTX/\beta CTX$ might be clin. important in diagnosing metabolic bone diseases.

ST type I collagen degrdn product age; urine collagen degrdn product age; telopeptide type I collagen degrdn age

IT Urine

(characterization of urinary degradation products derived from type I collagen)

IT Aging, animal

Development, mammalian postnatal

(characterization of urinary degradation products derived from type I collagen of young and old human bones)

IT Bone

(cortical; characterization of urinary degradation products derived from type I collagen of young and old human bones)

IT Bone

(trabecula; characterization of urinary degradation products derived from type I collagen of young and old human bones)

IT Peptides, biological studies

RL: BOC (Biological occurrence); BPR (Biological process); BSU (Biological study, unclassified); PRP (Properties); BIOL (Biological study); OCCU (Occurrence); PROC (Process)

(type I collagen C-terminal telopeptides; characterization of urinary degradation products derived from type I collagen of young and old human bones)

IT Collagens, biological studies

RL: BOC (Biological occurrence); BPR (Biological process); BSU (Biological study, unclassified); PRP (Properties); BIOL (Biological study); OCCU (Occurrence); PROC (Process)

(type I, C-terminal telopeptides; characterization of urinary degradation products derived from type I collagen of young and old human bones)

IT 3790-51-0 3790-52-1 63800-01-1, Hydroxylysylpyridinoline 87672-07-9 RL: BOC (Biological occurrence); BPR (Biological process); BSU (Biological study, unclassified); PRP (Properties); BIOL (Biological study); OCCU (Occurrence); PROC (Process)

(characterization of urinary degradation products derived from type I collagen of young and old human bones)

RE.CNT 41 THERE ARE 41 CITED REFERENCES AVAILABLE FOR THIS RECORD RE

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L81
     ANSWER 10 OF 29 HCAPLUS COPYRIGHT 2006 ACS on STN
ΑN
     1997:203828 HCAPLUS
DN
     126:183499
ED
     Entered STN: 28 Mar 1997
     Determination of type II-collagen
     telopeptide for bone disease diagnosis
     Nakamoto, Tadakatsu; Pponda, Hitomi; Kobayashi, Shinji; Hosoda, Kenji
ΙN
     Teijin Ltd, Japan
PA
     Jpn. Kokai Tokkyo Koho, 7 pp.
     CODEN: JKXXAF
DT
     Patent
LA
     Japanese
IC
     ICM
          G01N0033-53
          C07K0016-18; C12N0015-02; C12P0021-08; G01N0033-531; G01N0033-535;
          G01N0033-577; C12R0001-91
     9-2 (Biochemical Methods)
     Section cross-reference(s): 14
FAN.CNT 1
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PATENT NO.
                        KIND
                               DATE
                                         APPLICATION NO.
                                                                DATE
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     JP 09021803
                        A2
                                          JP 1995-172196
PΤ
                               19970121
                                                                19950707 <--
PRAI JP 1995-172196
                               19950707 <--
CLASS
 PATENT NO.
              CLASS PATENT FAMILY CLASSIFICATION CODES
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 JP 09021803
                ICM
                       G01N0033-53
                ICS
                       C07K0016-18; C12N0015-02; C12P0021-08; G01N0033-531;
                       G01N0033-535; G01N0033-577; C12R0001-91
                IPCI
                       G01N0033-53 [ICM, 6]; C07K0016-18 [ICS, 6]; C12N0015-02
                       [ICS, 6]; C12P0021-08 [ICS, 6]; G01N0033-531 [ICS, 6];
                       G01N0033-535 [ICS, 6]; G01N0033-577 [ICS, 6]; C12R0001-91
                       [ICS, 6]
    A simple and accurate method for detecting the type II
AB
     -collagen telopeptide in mammalian body fluids is described
     using monoclonal antibodies in the so-called sandwich method for
     the diagnosis of metabolism disorders of the cartilage. A kit for the
determination
    method is presented.
ST
     collagen telopeptide detn body fluid diagnosis
ΙT
     Bone, disease
        (determination of type II-collagen telopeptides
        for bone disease diagnosis)
IT
     Antibodies
     RL: ARG (Analytical reagent use); ANST (Analytical study); USES (Uses)
        (monoclonal; in determination of type II-collagen
        telopeptides for bone disease diagnosis)
     Peptides, analysis
IT
     RL: ANT (Analyte); ANST (Analytical study)
        (telo-; determination of type II-collagen
        telopeptides for bone disease diagnosis)
TT
     Collagens, analysis
     RL: ANT (Analyte); ANST (Analytical study)
        (type II; determination of type II-
        collagen telopeptide for bone disease diagnosis)
1.81
    ANSWER 11 OF 29 HCAPLUS COPYRIGHT 2006 ACS on STN
AN
    1997:81688 HCAPLUS
DN
    126:168697
ED
    Entered STN: 05 Feb 1997
TΤ
    Isomerized molecules in serum derived from bone resorption
ΑU
    Cloos, P. A. C.; Bonde, M.; Fledelius, C.; Christgau, S.;
     Christiansen, C.
CS
    Osteometer BioTech A/S, Herlev, DK-2730, Den.
SO
     International Congress Series (1996), 1118 (Osteoporosis 1996),
     227-231
     CODEN: EXMDA4; ISSN: 0531-5131
PB
    Elsevier
DT
    Journal
LA
    English
CC
     9-10 (Biochemical Methods)
     Section cross-reference(s): 1, 13, 14
    We developed 2 serum-based ELISAs, CrossLaps serum ELISA
AB
     (β-CLS) and α-CrossLaps serum ELISA (α-CLS),
    measuring the isopeptide and normal peptide form, resp., of the
     collagen type I specific sequence EKAHDGGR. In the isomerized
     form, the aspartyl residue (D) is linked to the glycine residue (G) via
     the \beta-carboxyl group of the side chain rather than through the
     \alpha-carboxyl group. The aim of the present study was to investigate
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the clin. importance of isomerization for the assessment of bone resorption in serum. The effect of bisphosphonate therapy on postmenopausal women was evaluated with the two ELISAs. While serum samples from all women treated with bisphosphonate displayed significant decreases after 9 mo of therapy when measured in the β -CLS assay (mean decrease \pm SEM, 56.8 \pm 4.0%), α -CLS values only decreased slightly (19.7 \pm 6.3%). It is suggested that isopeptides in serum recognized by the $\beta\text{-CLS}$ assay are derived from bone resorption, whereas the corresponding nonisomerized peptides (measured by α -CLS) also reflect metabolism of nonskeletal tissue. bone resorption serum peptide isomer ELISA; enzyme immunoassay peptide bone resorption; collagen peptide isomer ELISA bone resorption; postmenopause bisphosphonate therapy serum collagen peptide Blood analysis Urine analysis (ELISA of isomerized mols. in serum derived from bone resorption) Menopause (postmenopause; ELISA of isomerized mols. in serum derived from bone resorption) (resorption; ELISA of isomerized mols. in serum derived from bone resorption) Collagens, analysis RL: ANT (Analyte); THU (Therapeutic use); ANST (Analytical study); BIOL (Biological study); USES (Uses) (type I; ELISA of isomerized mols. in serum derived from bone resorption) 162929-64-8 187269-53-0 RL: ANT (Analyte); THU (Therapeutic use); ANST (Analytical study); BIOL (Biological study); USES (Uses) (ELISA of isomerized mols. in serum derived from bone resorption) 13598-36-2D, Phosphonic acid, alkylidenebis-derivs. RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses) (ELISA of isomerized mols. in serum derived from bone resorption) RE.CNT 10 THERE ARE 10 CITED REFERENCES AVAILABLE FOR THIS RECORD (1) Bonde, M; Clin Chem 1994, V40, P2022 HCAPLUS (2) Bonde, M; Clin Chem In press 1996 (3) Bonde, M; J Bone Min Res Abstract S481 1995, V10(Suppl 1) (4) Bonde, M; J Clin Endocrinol Metab 1995, V80, P864 HCAPLUS (5) Fledelius, C; J Biol Chem In press 1996 (6) Fledelius, C; J Bone Min Res Abstract S482 1995, V10(Suppl 1) (7) Garnero, P; J Bone Min Res Abstract 6 1995, V10(Suppl 1) (8) Hanson, D; J Bone Min Res 1992, V7, P1251 HCAPLUS (9) Kemp, P; Biochem J 1988, V252, P387 HCAPLUS (10) Wright, H; Crit Rev Biochem Mol Biol 1991, V26, P1 HCAPLUS L81 ANSWER 12 OF 29 HCAPLUS COPYRIGHT 2006 ACS on STN 1996:623921 HCAPLUS 125:296532 Entered STN: 21 Oct 1996 Coated-tube radioimmunoassay for C-telopeptides of type I collagen to assess bone resorption Bonde, Martin; Fledelius, Christian; Qvist, Per; Christiansen,

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Claus
CS
     Osteometer BioTech A/S, Herlev, DK-2730, Den.
SO
     Clinical Chemistry (Washington, D. C.) (1996), 42(10), 1639-1644
     CODEN: CLCHAU; ISSN: 0009-9147
PB
     American Association for Clinical Chemistry
DT
     Journal
LA
     English
CC
     9-10 (Biochemical Methods)
AΒ
     We present a coated-tube RIA that is useful for assessment of bone
     resorption. The assay uses a monoclonal antibody raised against a linear
     8-amino-acid sequence (EKAH-DGGR) derived from the C-telopeptides of type
     I collagen. Within-run and total CVs were 4.4% and 5.3-6.2%,
     resp., at concns. of 1-7 mg/L (n = 4-20). Anal. recovery was 98% \pm 8%
     and dilution 97% \pm 7%. Values obtained in a group of 36 premenopausal
     women were 227 \pm 89.6 mg/mol creatinine. In a group of 141
     postmenopausal women, the values obtained were 429 \pm 225 mg/mol
     creatinine, a highly significant increase of 89% (P < 0.001) over the
     premenopausal value. In a double-blind placebo-controlled clin. study of
     these postmenopausal women receiving five different doses of a
     bisphosphonate, a significant decrease of RIA-measured C-telopeptide
     values was seen in all bisphosphonate-treated groups, after just 3 mo.
     Values in urine samples from postmenopausal women assayed with the RIA (y)
     and the CrossLapsTM ELISA (x) agreed well: slope = 0.98 (95%
     confidence interval, 9.94-1.01), intercept = 0.34(0.25-0.43) mg/L, and
     Sylx = 0.93 mg/L (n = 678). We conclude that this RIA represents a
     valuable tool for assessing bone resorption.
     bone resorption RIA; immunoassay peptide collagen
ST
ΙT
     Bone
     Urine analysis
        (coated-tube RIA for C-telopeptides of type I collagen to
        assess bone resorption)
IT
     Peptides, analysis
     RL: ANT (Analyte); ANST (Analytical study)
        (coated-tube RIA for C-telopeptides of type I collagen to
        assess bone resorption)
IT
     Antibodies
     RL: ARG (Analytical reagent use); ANST (Analytical study); USES (Uses)
        (monoclonal, coated-tube RIA for C-telopeptides of type I
        collagen to assess bone resorption)
IT
     Collagens, biological studies
     RL: BSU (Biological study, unclassified); BIOL (Biological study)
        (type I, coated-tube RIA for C-telopeptides of type I collagen
        to assess bone resorption)
L81
     ANSWER 13 OF 29 HCAPLUS COPYRIGHT 2006 ACS on STN
ΑN
     1996:359859 HCAPLUS
DN
     125:29628
     Entered STN: 21 Jun 1996
ED
     Estimation of the fragmentation pattern of collagen in body
     fluids and the diagnosis of disorders associated with the metabolism of
     collagen
     Bonde, Martin; Qvist, Per
IN
PΑ
     Osteometer Bio Tech A/s, Den.
     PCT Int. Appl., 40 pp.
SO
     CODEN: PIXXD2
DT
     Patent
LA
     English
IC
     ICM G01N0033-68
     ICS C07K0007-06
     9-16 (Biochemical Methods)
CC
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gitomer - 10 / 730070 Section cross-reference(s): 14 FAN.CNT 2 PATENT NO. APPLICATION NO. KIND DATE -----______ -----____ -----A1 19960425 WO 1995-EP4055 19951016 <--WO 9612193 PΙ W: AL, AM, AT, AU, BB, BG, BR, BY, CA, CH, CN, CZ, DE, DK, EE, ES, FI, GB, GE, HU, IS, JP, KE, KG, KP, KR, KZ, LK, LR, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, TJ RW: KE, MW, SD, SZ, UG, AT, BE, CH, DE, DK, ES, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, ML, MR, NE, SN, TD, TG AU 9537462 A1 19960506 AU 1995-37462 19951016 <--EP 1995-935451 A1 19951016 <--EP 787301 19970806 EP 787301 B1 20010214 R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IE, IT, LI, LU, MC, NL, PT, SE JP 10507266 T2 19980714 JP 1995-512947 19951016 <--AT 199185 Ε 20010215 AT 1995-935451 19951016 <--20010416 ES 1995-935451 19961003 WO 1996-EP1228 ES 2154739 Т3 19951016 <--WO 9630765 A1 19960321 <--W: AL, AM, AT, AU, AZ, BB, BG, BR, BY, CA, CH, CN, CZ, DE, DK, EE, ES, FI, GB, GE, HU, IS, JP, KE, KG, KP, KR, KZ, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI RW: KE, LS, MW, SD, SZ, UG, AT, BE, CH, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN AU 9651468 A1 19961016 AU 1996-51468 19960321 <--AU 712375 B2 19991104 **A1** EP 820598 19980128 EP 1996-908100 19960321 <--R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, FI CN 1185209 19980617 CN 1996-194134 Α 19960321 <--A 19980630 T2 19990302 B1 20010403 A 20000829 BR 9607854 Α BR 1996-7854 19960321 <--JP 11502622 Т2 JP 1996-528893 19960321 <--B1 20010403 US 1997-817397
US 6110689 A 20000829 US 1997-963825
US 6323314 B1 20011127 US 2000-500811
US 6355442 B1 20020312 US 2000-548608
US 6342361 B1 20020129 US 2000-570573
US 6372442 B1 20020416 US 2000-714146
US 2003119058 A1 20030626 US 2002-58124

PRAI DK 1994-1194 A 19941017 <-
GB 1995-6050 A 19950324 <-US 1994-187319 B1 19940121 <-WO 1995-EP4055 W 19951016 <-WO 1996-EP1228 W 19960321 <--19970611 <--19971104 <--20000210 <--20000413 <--20000512 <--US 2000-714146 20001117 <--20020129 <--A1 US 1997-817397 19970611 <--A1 US 1997-963825 19971104 <--US 2000-570573 A3 20000512 CLASS PATENT NO. CLASS PATENT FAMILY CLASSIFICATION CODES WO 9612193 ICM G01N0033-68 ICS C07K0007-06 IPCI G01N0033-68 [ICM, 6]; C07K0007-06 [ICS, 6]; C07K0007-00

CLASS PATENT FAMILY CLASSIFICATION CODES

ICM G01N0033-68
ICS C07K0007-06
IPCI G01N0033-68 [ICM,6]; C07K0007-06 [ICS,6]; C07K0007-00
[ICS,6]
IPCR C07K0014-435 [I,C]; C07K0014-78 [I,A]; C07K0016-18
[I,A]; C07K0016-18 [I,C]; G01N0033-68 [I,A];
G01N0033-68 [I,C]
ECLA C07K014/78; C07K016/18; G01N033/68R

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EP 787301
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JP 10507266
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AU 9651468
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                        C07K0016-18 [ICS, 6]; C12N0005-20 [ICS, 6]
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EP 820598
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                        C07K0016-18 [ICS, 6]; C12N0005-20 [ICS, 6]
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CN 1185209
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                        C07K0016-18 [ICS, 6]; C12N0005-20 [ICS, 6]
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BR 9607854
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JP 11502622
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                NCL
                        435/007.100; 435/007.920; 435/007.930; 435/007.940;
                        435/007.950; 436/518.000; 436/532.000; 530/356.000;
                        530/388.100; 530/389.100
                ECLA
                       C07K014/78; G01N033/68R
                       G01N0033-53 [ICM, 7]
US 6110689
                IPCI
                IPCR
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[I,A]; G01N0033-68 [I,C]
                 NCL
                        435/007.100; 435/007.930; 435/007.940; 435/070.210;
                        435/331.000; 435/975.000; 436/518.000; 436/536.000;
                        436/548.000; 436/815.000; 530/300.000; 530/323.000;
                        530/328.000; 530/387.900; 530/388.100; 530/391.100;
                        530/391.300
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                        C07K016/18; G01N033/68R
 US 6323314
                 IPCI
                        A61K0038-04 [ICM, 7]
                 IPCR
                        C07K0016-18 [I,A]; C07K0016-18 [I,C]; G01N0033-68
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                        530/356,000
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                        C07K016/18; G01N033/68R
 US 6355442
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                        G01N0033-53 [ICM, 7]
                 IPCR
                        C07K0016-18 [I,A]; C07K0016-18 [I,C]; G01N0033-68
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                        530/328.000; 530/356.000; 530/387.900; 530/388.100
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                        435/007.100; 435/007.920; 435/007.930; 435/975.000;
                        436/518.000; 436/531.000; 530/328.000; 530/329.000;
                        530/330.000; 530/331.000; 530/356.000; 530/387.900;
                        530/388.100
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 US 6372442
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                        [I,A]; G01N0033-68 [I,C]
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                        G01N0033-543 [ICS,7]; C07K0002-00 [ICS,7]; C07K0004-00
                        [ICS,7]; C07K0005-00 [ICS,7]; C07K0007-00 [ICS,7];
                        C07K0014-00 [ICS,7]; C07K0016-00 [ICS,7]; C07K0017-00
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                        C07K0001-00 [ICS,7]; C09H0001-00 [ICS,7]; A61K0038-17
                        [ICS, 7]; G01N0033-545 [ICS, 7]; G01N0033-544 [ICS, 7]
                 IPCR
                        C07K0016-18 [I,A]; C07K0016-18 [I,C]; G01N0033-68
                        [I,A]; G01N0033-68 [I,C]
                 NCL
                        435/007.100
                 ECLA
                        C07K016/18; G01N033/68R
AB
     The fragmentation pattern of collagen, especially of type 1, as
     reflected in breakdown products of collagen in a body fluid such
     as serum or urine is estimated by measuring the levels of such breakdown
     products using two or more distinct immunoassays. The results may be
     combined into a numerical index diagnostic of one or more pathol.
     conditions or patient types.
ST
     collagen body fluid diagnosis disorder metab
TT
     Blood analysis
     Body fluid
     Diagnosis
     Immunoassay
     Urine analysis
        (estimation of the fragmentation pattern of collagen in body
        fluids and the diagnosis of disorders associated with the metabolism of
        collagen)
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ΙT
     Collagens, analysis
     RL: ANT (Analyte); ANST (Analytical study)
        (estimation of the fragmentation pattern of collagen in body
        fluids and the diagnosis of disorders associated with the metabolism of
        collagen)
ΙT
     Collagens, analysis
     RL: ANT (Analyte); ANST (Analytical study)
        (type I, estimation of the fragmentation pattern of collagen in
        body fluids and the diagnosis of disorders associated with the metabolism of
        collagen)
L81 ANSWER 14 OF 29 HCAPLUS COPYRIGHT 2006 ACS on STN
AN
     1995:616314 HCAPLUS
     123:161872
DN
ED
     Entered STN: 16 Jun 1995
TΙ
     The detection of the mRNAs of procollagen types I,
     II and III in human fetal fingers by in situ hybridization using
     digoxigenin-labeled oligonucleotide probes
ΑU
     Hamada, K.; Okawara, Y.; Fryer, J. N.; Tomonaga, A.; Fukuda, H.
     Oiso Hospital, Tokai University, Kanagawa, 259-01, Japan
CS
     Histochemical Journal (1995), 27(4), 309-17
SO
     CODEN: HISJAE; ISSN: 0018-2214
DT
     Journal
     English
LA
CC
     3-1 (Biochemical Genetics)
     Section cross-reference(s): 9, 13
AB
     The mRNAs encoding procollagen \alpha1
                                       type I,
        type II and αl type III have
     been localized in paraffin sections of human fetal fingers using
     digoxigenin-labeled synthetic oligonucleotide probes. The probe-mRNA
     hybrids were visualized using an anti-digoxin antibody amplified with
     sandwich techniques. These protocols provided an excellent
     hybridization signal with minimal background noise. The sensitivity of
     the protocols was nearly equivalent to that seen when using isotopic cDNA
     probes. In human fetal fingers, intense hybridization signals for
     procollagen \alpha1 type I mRNA were detected in the osteoblasts
     and the fibroblasts of periosteum and perichondrium, the tenocytes of
     tendons, fibroblasts of ligaments, the synovial membrane and deeper layers
     of the dermis. In contrast, pos. hybridization signals for
     procollagen α1 type II mRNA were
     visualized in chondrocytes and the cambial layer of perichondrium.
     signals for procollagen al type III mRNA were detected in
     the fibroblasts of the dermis and perichondrium. The probes which have
     lower melting temps. (Tm) could not detect the corresponding mRNAs.
ST
     mRNA procollagen human fetus digoxigenin probe; hybridization
     digoxigenin oligonucleotide probe
IT
     Ligament
     Synovial membrane
        (detection of procollagen mRNA; detection of mRNAs of
        procollagen types I, II and III in human
        fetal fingers by in situ hybridization using digoxigenin-labeled
        oligonucleotide probes)
ΤT
     Ribonucleic acids, messenger
     RL: BOC (Biological occurrence); BSU (Biological study, unclassified);
     BIOL (Biological study); OCCU (Occurrence)
        (detection; detection of mRNAs of procollagen types
        I, II and III in human fetal fingers by in situ hybridization
        using digoxigenin-labeled oligonucleotide probes)
IT
     Nucleic acid hybridization
        (in situ, non-isotopic; detection of mRNAs of procollagen
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types I, II and III in human fetal fingers by in situ
        hybridization using digoxigenin-labeled oligonucleotide probes)
ΙT
     Cartilage
        (perichondrium, detection of procollagen mRNA; detection of
        mRNAs of procollagen types I, II and III
        in human fetal fingers by in situ hybridization using
        digoxigenin-labeled oligonucleotide probes)
IT
     Bone
        (periosteum, detection of procollagen mRNA; detection of
        mRNAs of procollagen types I, II and III
        in human fetal fingers by in situ hybridization using
        digoxigenin-labeled oligonucleotide probes)
IT
    Collagens, biological studies
     RL: BOC (Biological occurrence); BSU (Biological study, unclassified); BUU
     (Biological use, unclassified); BIOL (Biological study); OCCU
     (Occurrence); USES (Uses)
        (pro-, αl of type I, II and III; detection of
        mRNAs of procollagen types I, II and III
        in human fetal fingers by in situ hybridization using
        digoxigenin-labeled oligonucleotide probes)
ΙT
        (tenocyte, detection of procollagen mRNA; detection of mRNAs
        of procollagen types I, II and III in
        human fetal fingers by in situ hybridization using digoxigenin-labeled
        oligonucleotide probes)
L81
   ANSWER 15 OF 29 HCAPLUS COPYRIGHT 2006 ACS on STN
ΑN
    1995:541494 HCAPLUS
DN
    122:286078
ED
    Entered STN: 11 May 1995
TT
    A method of assaying collagen fragments in body fluids, a test
    kit and means for carrying out the method and use of the method to
    diagnose the presence of disorders associated with the metabolism of
    collagen
ΙN
    Qvist, Per; Bonde, Martin
PA
    Osteometer A/S, Den.
SO
    PCT Int. Appl., 87 pp.
    CODEN: PIXXD2
DΤ
    Patent
LA
    English
IC
    ICM G01N0033-53
    ICS G01N0033-68; C07K0014-78
CC
    9-10 (Biochemical Methods)
FAN.CNT 1
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                               DATE
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                                         WO 1994-DK348
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                              19950323
                         A1
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            MN, MW, NL, NO, NZ, PL, PT, RO, RU, SD, SE, SI, SK, TJ, TT, UA,
            US, UZ
         RW: KE, MW, SD, AT, BE, CH, DE, DK, ES, FR, GB, GR, IE, IT, LU, MC,
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                         A1
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                                           JP 1995-508839
                                                                  19940919 <--
    JP 3423720
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                               20010606
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PRAI DK 1993-1040
                         Α
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     WO 1994-DK348
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                                19940919
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CLASS
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                 CLASS PATENT FAMILY CLASSIFICATION CODES
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 WO 9508115
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                        [I,A]; G01N0033-68 [I,C]
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                        C07K014/78; G01N033/68R
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                        [I,A]; G01N0033-68 [I,C]
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                 IPCI
                        G01N0033-53 [ICM,7]; C07K0005-083 [ICS,7]; C07K0005-00
                        [ICS,7]; C07K0007-06 [ICS,7]; C07K0007-00 [ICS,7];
                        C07K0014-78 [ICS,7]; C07K0014-435 [ICS,7]
 EP 1104887
                 IPCI
                        G01N0033-68 [ICM, 6]; C07K0014-78 [ICS, 6]; C07K0014-435
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                 ECLA
                        G01N033/68R
 AT 209356
                 IPCI
                        G01N0033-53 [ICM,7]; G01N0033-68 [ICS,7]; C07K0014-78
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                        C07K0014-435 [I,C]; C07K0014-78 [I,A]; G01N0033-68
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 JP 2003202338
                 IPCI
                        G01N0033-53 [ICM,7]; C07K0016-18 [ICS,7]; C12N0005-10
                        [ICS,7]; G01N0033-531 [ICS,7]; C12N0015-02 [ICS,7];
                        C12P0021-08 [ICS, 7]
     A method of assaying collagen fragments in body fluids(such as
AB
     urine, blood), including bringing a sample of body fluid in contact with
     at least one immunol. binding partner for the collagen
     fragments, said binding partner being immunoreactive with synthetic
     peptides, the sequences of which are essentially derived from
     collagen and containing potential sites for crosslinking. The
     immunol. binding partners are incorporated, either as whole antibodies or
     as immunol. active fragments thereof, in an assay for quant. determination of
     collagen fragments in the sample. In addition to being contacted
     with the immunol. binding pattern(s), the sample may be thought into
     direct contact with the corresponding peptide. The invention further
     comprise a test kit and specific means for carrying out the method.
     structure of specific peptides is also described.
ST
     body fluid collagen fragment ELISA; immunoassay bone
     collagen antibody peptide sequence
ΙT
     Animal cell line
     Blood analysis
     Body fluid
     Bone
     Synovial fluid
     Urine analysis
        (a method of assaying collagen fragments in body fluids, a
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test kit and means for carrying out the method and use of the method to diagnose the presence of disorders associated with the metabolism of collagen) IT Collagens, biological studies RL: BOC (Biological occurrence); BSU (Biological study, unclassified); BIOL (Biological study); OCCU (Occurrence) (a method of assaying collagen fragments in body fluids, a test kit and means for carrying out the method and use of the method to diagnose the presence of disorders associated with the metabolism of collagen) TT Antibodies RL: BPR (Biological process); BSU (Biological study, unclassified); BUU (Biological use, unclassified); BIOL (Biological study); PROC (Process); USES (Uses) (a method of assaying collagen fragments in body fluids, a test kit and means for carrying out the method and use of the method to diagnose the presence of disorders associated with the metabolism of collagen) ΙT Immunoassay (enzyme-linked immunosorbent assay, a method of assaying collagen fragments in body fluids, a test kit and means for carrying out the method and use of the method to diagnose the presence of disorders associated with the metabolism of collagen) ΙT 83462-55-9, Deoxypyridinoline RL: ANT (Analyte); ANST (Analytical study) (a method of assaying collagen fragments in body fluids, a test kit and means for carrying out the method and use of the method to diagnose the presence of disorders associated with the metabolism of collagen) IT 71227-72-0 146663-75-4 162929-64-8 162929-65-9 162929-66-0 162929-67-1 162929-69-3 162929-68-2 162929-70-6 162929-71-7 162929-73-9 162929-72-8 162929-74-0 162929-75-1 162929-76-2 162929-77-3 162929-78-4 RL: BUU (Biological use, unclassified); BIOL (Biological study); USES (Uses) (a method of assaying collagen fragments in body fluids, a test kit and means for carrying out the method and use of the method to diagnose the presence of disorders associated with the metabolism of collagen) L81 ANSWER 16 OF 29 HCAPLUS COPYRIGHT 2006 ACS on STN ΑN 1995:437464 HCAPLUS 122:209095 DN ED Entered STN: 23 Mar 1995 Applications of an enzyme immunoassay for a new marker of bone resorption (CrossLaps): follow-up on hormone replacement therapy and osteoporosis risk assessment ΑU Bonde, Martin; Qvist, Per; Fledelius, Christian; Riis, Bente Juel; Christiansen, Claus CS Center for Clinical and Basic Research, Ballerup, DK-2750, Den. SO Journal of Clinical Endocrinology and Metabolism (1995), 80(3), CODEN: JCEMAZ; ISSN: 0021-972X PB Endocrine Society DT Journal LA English

An enzyme-linked immunosorbent immunoassay (ELISA) for a new

CC

AΒ

9-10 (Biochemical Methods)

Section cross-reference(s): 2, 14

marker of bone resorption (CrossLaps) and evaluated. The ELISA procedure dets. degradation products of type I collagen in urine. Values obtained in the ELISA and in pyridinoline by high pressure liquid chromatog. were correlated after a correction for creatinine. A high correlation was found (r = 0.77). A group of postmenopausal women showed an increase of more than 70% compared to values in premenopausal women. Hydroxyproline was increased by 23%, osteocalcin by 52%, pyridinoline by 31%, and deoxypyridinoline by 50%. A highly significant decrease (60.7%) in the CrossLaps values was seen after 12 mo in samples from patients receiving hormone replacement therapy compared to a placebo group. The spontaneous bone loss in an untreated group of women was determined by repeated forearm bone mass measurement over 24 mo. Baseline values obtained in the CrossLaps ELISA were correlated to the rate of loss, yielding a highly significant r value of -0.61, indicating that CrossLaps might be a useful parameter for assessment of the risk of osteoporosis in postmenopausal women. enzyme immunoassay marker bone resorption; hormone replacement therapy osteoporosis risk Bone Osteoporosis Urine analysis (applications of enzyme immunoassay for a new marker of bone resorption (CrossLaps): follow-up on hormone replacement therapy and osteoporosis risk assessment) Hormones RL: ANT (Analyte); ANST (Analytical study) (applications of enzyme immunoassay for a new marker of bone resorption (CrossLaps): follow-up on hormone replacement therapy and osteoporosis risk assessment) Biological transport (resorption; applications of enzyme immunoassay for a new marker of bone resorption (CrossLaps): follow-up on hormone replacement therapy and osteoporosis risk assessment) Collagens, analysis RL: ANT (Analyte); ANST (Analytical study) (type I, degradation products; applications of enzyme immunoassay for a new marker of bone resorption (CrossLaps): follow-up on hormone replacement therapy and osteoporosis risk assessment) L81 ANSWER 17 OF 29 HCAPLUS COPYRIGHT 2006 ACS on STN 1995:257916 HCAPLUS 122:27271 Entered STN: 22 Dec 1994 Sandwich immunoassay for collagen Amano, Satoshi; Masuda, Yoshiko; Ito, Shiqeki; Fujio, Mieko Shiseido Co., Ltd., Japan; Nippon Shoji KK Jpn. Kokai Tokkyo Koho, 10 pp. CODEN: JKXXAF Patent Japanese ICM G01N0033-53 ICS G01N0033-577 ICA A61B0010-00 9-10 (Biochemical Methods) FAN.CNT 1 PATENT NO. DATE APPLICATION NO. DATE KIND ____ ---------------

ST

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CLASS

JP 06242109

PRAI JP 1991-80891

A2

JP 1991-80891

19910318 <--

19940902

19910318 <--

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CLASS PATENT FAMILY CLASSIFICATION CODES
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 JP 06242109
                ICM
                       G01N0033-53
                ICS
                       G01N0033-577
                ICA
                       A61B0010-00
                IPCI
                       G01N0033-53 [ICM, 5]; G01N0033-577 [ICS, 5]; A61B0010-00
                       [ICA, 5]
AΒ
    Disclosed is a method comprising a pretreatment procedure of sample at
     39-60° and sandwich immunoassay with solid
    phase-immobilized and labeled monoclonal antibodies. The sensitivity of
     the immunoassay is largely increased and is used for determination of
     collagen, especially human collagen IV in blood serum, and for
    diagnosis of liver diseases, liver cancer, cirrhosis, etc.
ST
     sandwich immunoassay collagen liver disease
ΙT
    Blood analysis
    Cirrhosis
    Liver, disease
    Liver, neoplasm
        (sample pretreatment at 39-60° and sandwich
        immunoassay for collagen determination in blood serum and for
        diagnosis of liver diseases)
IT
    Collagens, analysis
    RL: ANT (Analyte); ANST (Analytical study)
        (sample pretreatment at 39-60° and sandwich
        immunoassay for collagen determination in blood serum and for
        diagnosis of liver diseases)
IT
    Antibodies
    RL: ARG (Analytical reagent use); THU (Therapeutic use); ANST (Analytical
     study); BIOL (Biological study); USES (Uses)
        (monoclonal, immobilized and labeled; sample pretreatment at
        39-60° and sandwich immunoassay for collagen
        determination in blood serum and for diagnosis of liver diseases)
ΙT
    Collagens, analysis
    RL: ANT (Analyte); ANST (Analytical study)
        (type IV, sample pretreatment at 39-60° and sandwich
        immunoassay for collagen determination in blood serum and for
        diagnosis of liver diseases)
L81
    ANSWER 18 OF 29 HCAPLUS COPYRIGHT 2006 ACS on STN
AN
    1994:696476 HCAPLUS
DN
    121:296476
ED
    Entered STN: 24 Dec 1994
    Immunoassay for quantifying type I collagen degradation products
TΤ
     in urine evaluated
ΑU
    Bonde, Martin; Qvist, Per; Fledelius, Christian; Riis, Bente
    Juel; Christiansen, Claus
CS
    Osteometer A/S, Rodovre, DK-2610, Den.
SO
    Clinical Chemistry (Washington, D. C.) (1994), 40(11, Pt. 1),
    2022-5
    CODEN: CLCHAU; ISSN: 0009-9147
PB
    American Association for Clinical Chemistry
DT
    Journal
LA
    English
CC
    9-10 (Biochemical Methods)
    Section cross-reference(s): 13
AΒ
    An ELISA for measuring type I collagen degradation
    products in urine <3 h was evaluated. The measuring range was 0.5-10.5
    mg/L with a detection limit of 0.2 mg/L. Within-run and total CVs were
    5.3% and 6.6% resp. Anal. recovery averaged 100%. The mean (± SD)
    concns. in urine samples from a healthy premenopausal population (n = 102)
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were 250 \pm 110 mg/mol creatinine (Cr). A group of healthy
    postmenopausal women (n = 410) gave a mean value of 416 ± 189 mg/mol
    Cr. Values obtained in the ELISA correlated well (r = 0.83) to
    HPLC values for the established bone resorption marker deoxypyridinoline
    (n = 214), slightly better than the correlation to hydroxyproline
    measurements (r = 0.78, n = 421). We conclude that the assay described
    here presents a useful tool or further elucidating the importance of type
    I collagen degradation products in urine.
    immunoassay collagen degrdn product urine
    Menopause
    Urine analysis
        (immunoassay for quantifying type I collagen degradation products
       in urine evaluated)
    Immunoassay
       (enzyme-linked immunosorbent
       assay, immunoassay for quantifying type I collagen
       degradation products in urine evaluated)
    Collagens, analysis
    RL: ANT (Analyte); ANST (Analytical study)
       (type I, degradation products; immunoassay for quantifying type I
       collagen degradation products in urine evaluated)
L81 ANSWER 19 OF 29 HCAPLUS COPYRIGHT 2006 ACS on STN
    1993:76619 HCAPLUS
    118:76619
    Entered STN: 02 Mar 1993
    High-reproducibility sandwich immunoassay for collagen
    determination in serum
    Ito, Shigeki; Fujio, Mieko
    Nippon Shoji Co., Ltd., Japan
    Jpn. Kokai Tokkyo Koho, 8 pp.
    CODEN: JKXXAF
    Patent
    Japanese
    ICM G01N0033-53
    9-10 (Biochemical Methods)
FAN.CNT 1
                      KIND DATE APPLICATION NO. DATE
    PATENT NO.
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    JP 04324357
                      A2 19921113 JP 1991-94128 19910424 <--
PRAI JP 1991-94128
                             19910424 <--
CLASS
PATENT NO.
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JP 04324357
              ICM G01N0033-53
               IPCI G01N0033-53 [ICM, 5]
    In determining collagen in a sample by sandwich immunoassay
    (EIA), the collagen in the sample in incubated at 39-60°
    in the presence of anionic surfactants (SDS, Na lauryl benzenesulfonates
    or lithium laurylsulfate) to improve the anal. sensitivity and
    reproducibility. Determination of collagen in serum for diagnosis of
    chronic hepatitis and cirrhosis is given as an example. The patients
    showed elevated serum type IV collagen levels.
    collagen sandwich EIA anionic surfactant; heating
    surfactant EIA collagen hepatitis cirrhosis
    Firing, heat-treating process
       (anionic surfactant and, in collagen determination by
       sandwich EIA, to improve sensitivity and reproducibility)
    Collagens, analysis
    RL: ANST (Analytical study)
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(chemical of, in serum, by sandwich EIA, anionic surfactants
        for)
IT
     Blood analysis
        (collagen determination in, by sandwich EIA, anionic
        surfactants for)
IT
     Cirrhosis
        (diagnosis of, type IV collagen determination in serum for)
TΤ
     Surfactants
        (anionic, heat-treatment and, in collagen determination by
        sandwich EIA, to improve sensitivity and reproducibility)
IT
     Hepatitis
        (chronic, diagnosis of, type IV collagen determination in serum for)
     Immunoassay
TΨ
        (enzyme, sandwich, collagen determination by, anionic
        surfactants for)
ΙT
     Collagens, analysis
     RL: ANST (Analytical study)
        (type IV, chemical of, in serum, by sandwich EIA, anionic
        surfactants for)
IT
     151-21-3, SDS, uses
                          2044-56-6, Lithium laurylsulfate 25155-30-0,
     Sodium laurylbenzenesulfonate
     RL: USES (Uses)
        (in collagen determination by sandwich EIA, to improve
        sensitivity and reproducibility)
L81
    ANSWER 20 OF 29 HCAPLUS COPYRIGHT 2006 ACS on STN
AN
     1993:3437 HCAPLUS
DN
    118:3437
    Entered STN: 10 Jan 1993
ΕD
TΙ
    Kit for collagen determination
IN
    Amano, Satoshi; Masuda, Yoshiko; Yoshida, Tsuyoshi; Asamatsu, Chinatsu;
    Itoh, Shigeki
PA
    Shiseido Co., Ltd., Japan; Nipponshoji Co., Ltd.
SO
    PCT Int. Appl., 66 pp.
    CODEN: PIXXD2
DT
    Patent
LA
    Japanese
    ICM G01N0033-577
ICS G01N0033-53; G01N0033-531
     9-15 (Biochemical Methods)
    Section cross-reference(s): 14
FAN.CNT 1
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                        KIND
                                         APPLICATION NO. DATE
                               DATE
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PΙ
    WO 9216846
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                               19921001 WO 1992-JP328
                                                                 19920318 <--
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        RW: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LU, MC, NL, SE
                A2 19921014 JP 1991-80892
    JP 04289455
                                                                19910318 <--
    EP 535239
                        A1
                               19930407
                                           EP 1992-906928
                                                                 19920318 <--
        R: DE, FR, GB, IT, NL
    JP 05209883
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                                        JP 1992-232726 19920806 <--
PRAI JP 1991-80892
                        Α
                               19910318 <--
    JP 1991-237134
WO 1992-JP328
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WO 9216846
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                ICS
                       G01N0033-53; G01N0033-531
                IPCI
                       G01N0033-577 [ICM, 5]; G01N0033-53 [ICS, 5]; G01N0033-531
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[ICS, 5]
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                        G01N0033-68 [I,A]; G01N0033-68 [I,C]
 JP 04289455
                 IPCI
                        G01N0033-53 [ICM,5]; A61B0010-00 [ICS,5]; G01N0033-577
                        [ICS, 5]
                 IPCI
                        G01N0033-577 [ICM,5]; G01N0033-53 [ICS,5]; G01N0033-531
 EP 535239
                        [ICS, 5]
                        G01N0033-68 [I,A]; G01N0033-68 [I,C]
                 IPCR
 JP 05209883
                        G01N0033-531 [ICM,5]; G01N0033-53 [ICS,5]; G01N0033-577
                 IPCI
                        [ICS, 5]
     Chaotropic agents, heparins, chelating agents, gelatins, or albumins are
AΒ
     used to enhance the sensitivity of enzymic immunoanal. (EIA) for
     collagen in serum sample. Monoclonal antibody to collagen
     IV was raised and used in sandwich-type immunoassay (in the
     presence of gelatin, sodium heparin, EDTA, and sodium thiocyanate) for
     diagnosis of chronic hepatitis and hepatocirrhosis.
ST
     collagen detn monoclonal antibody
ΙT
     Blood analysis
        (collagen determination in, enhancer for EIA for)
IT
     Chelating agents
     Gelatins, uses
     RL: USES (Uses)
        (immunoassay of collagen with)
TT
     Collagens, analysis
     RL: ANST (Analytical study)
        (immunoassay of, enhancer for)
ΙT
     Fibronectins
     Laminins
     RL: ANST (Analytical study)
        (monoclonal antibody to, in EIA, for collagen determination)
TΤ
     Denaturants
        (chaotropic, immunoassay of collagen with)
     Albumins, compounds
ፐጥ
     RL: ANST (Analytical study)
        (compds., immunoassay of collagen with)
TΤ
     Antibodies
     RL: ANST (Analytical study)
        (monoclonal, to collagen, in immunoassay)
TΤ
     Collagens, analysis
     RL: ANST (Analytical study)
        (type I, immunoassay of)
IT
     Collagens, analysis
     RL: ANST (Analytical study)
        (type III, monoclonal antibody to, in EIA, for collagen
        determination)
ΙT
     Collagens, analysis
     RL: ANST (Analytical study)
        (type IV, monoclonal antibody to, in EIA, for collagen determination)
ΙT
     Collagens, analysis
     RL: ANST (Analytical study)
        (type V, monoclonal antibody to, in EIA, for collagen determination)
IT
     54-21-7, Sodium salicylate
                                 76-03-9, Trichloroacetic acid, biological
     studies
               4264-83-9
                           7681-11-0, Potassium iodide, biological studies
     9005-49-6, Heparin, biological studies
     RL: ANST (Analytical study)
        (collagen determination with)
IT
     9041-08-1, Sodium heparin 12678-07-8, Chondroitin sulfate C sodium salt
     37319-17-8, Pentosan polysulfate sodium 39422-86-1, Dextran sulfate
     potassium salt
                      39455-18-0, Chondroitin sulfate A sodium salt
     EDTA, biological studies
                               67-42-5, EGTA
                                               7447-40-7, Potassium chloride,
                          7601-89-0, Sodium perchlorate
     biological studies
                                                           7631-99-4, Sodium
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nitrate, biological studies
                                   7758-02-3, Potassium bromide, biological
     studies
     RL: ANST (Analytical study)
        (immunoassay of collagen with)
T.81
     ANSWER 21 OF 29 HCAPLUS COPYRIGHT 2006 ACS on STN
ΑN
     1991:629046 HCAPLUS
DN
     115:229046
ED
     Entered STN: 29 Nov 1991
ΤI
     Studies on secretion of surfactant protein A(SP-A) and D(SP-D) from
     alveolar type II cells
ΑU
     Miyamura, Kazuo; Ogasawara, Yoshinori; Kuroki, Yoshio
     Dep. Intern. Med., Sapporo Med. Coll., Sapporo, Japan
CS
     Sapporo Igaku Zasshi (1991), 60(2), 183-96
SO
     CODEN: SIZSAR; ISSN: 0036-472X
DT
     Journal
LA
     Japanese
CC
     13-2 (Mammalian Biochemistry)
     Section cross-reference(s): 9
AR
     Secretion of SP-A and SP-D from isolated alveolar type
     II cells and their subcellular distribution were studied.
     study secretion of SP-A in culture, a sensitive microassay for rat SP-A
     was developed, the secretion of SP-A and SP-D, both of which are
     characterized by collagen-like sequences and carbohydrate
     binding property, by primary cultures of rat alveolar type
     II cells was examined A sensitive sandwich ELISA
     was established using anti-rat SP-A Fab'-HRP conjugates. The assay system
     was capable of detecting SP-A as low as 0.1 ng/mL. Freshly isolated
     type II cells contained 21.91 ng SP-A/µg DNA and 5.05
     ng SP-D/\mug DNA. When type II cells were cultivated
     at 37° for 20 h, the intracellular content of SP-D decreased by
     .apprx.52.7%, whereas SP-A content increased 26.5%. During the
     cultivation of type II cells at 4° for 20 h,
     .apprx.3.05 ng SP-A/\mug DNA and 0.33 ng SP-D/\mug DNA, corresponding to
     .apprx.40% and 5% of the 37° secretion, resp., appeared to be
     secreted into the media. Secretions of SP-A, SP-D, and phospholipids were
     stimulated by TPA (10-7 M) and inhibited by ConA (10 \mu g/mL). The
     secretions of SP-A and SP-D stimulated with TPA were .apprx.160% compared
     with basal secretion, although phosphatidylcholine secretion was
     stimulated by .apprx.1300% by day 1 type II cells.
     Two hydrophilic surfactant proteins and disatd. phosphatidylcholine
     appeared to distribute in fractions with different densities obtained by
     discontinuous sucrose d. gradient centrifugation of type
     II cell homogenates. Thus, the metabolism and secretion of SP-A and
     SP-D appear to be regulated independently in type II
     cells, and SP-A and SP-D may be secreted through other pathways besides
     lamellar bodies.
ST
     surfactant protein secretion lung alveolus; ELISA surfactant
     protein
     Phosphatidylcholines, biological studies
     Phospholipids, biological studies
     RL: BIOL (Biological study)
        (secretion of, by lung type II cells, surfactant
        protein secretion in relation to)
ΙT
     Immunochemical analysis
        (enzyme-linked immunosorbent assay, of surfactant proteins)
TΤ
     Lung, metabolism
        (great alveolar cell, surfactant protein secretion by)
TT
     Proteins, specific or class
     RL: BIOL (Biological study)
```

(hydrophilic, surfactant, secretion of, by lung type II cells) IT Sialoglycoproteins RL: PROC (Process) (pulmonary surfactant-associated, SP-A (surfactant protein A), secretion of, by lung type II cells) IT Glycoproteins, specific or class RL: PROC (Process) (pulmonary surfactant-associated, SP-D (surfactant protein D), secretion of, by lung type II cells) IT 11028-71-0, Concanavalin A 16561-29-8, TPA RL: BIOL (Biological study) (surfactant protein secretion by lung type II cells in response to) L81 ANSWER 22 OF 29 HCAPLUS COPYRIGHT 2006 ACS on STN ΑN 1991:99419 HCAPLUS 114:99419 DN ED Entered STN: 23 Mar 1991 ΤI Serum level of vascular basement membrane associated collagen by the sandwich ELISA with monoclonal antibodies and its clinical significance in various diseases AU Tomura, Shigeo; Yoshida, Tsuyoshi; Shiba, Kiyoko; Kino, Jun; Cho, Hiroko; Asamatsu, Chinatsu; Nakajima, Keisuke; Miyake, Kazuhiko; Hayashi, Toshihiko CS Dep. Intern. Med., Nakano Gen. Hosp., Tokyo, 164, Japan SO Rinsho Byori (1990), 38(11), 1279-85 CODEN: RBYOAI; ISSN: 0047-1860 DT Journal LA Japanese CC 14-15 (Mammalian Pathological Biochemistry) A sandwich ELISA system for detecting vascular AB basement membrane associated collagen (BAC) was developed. Serum levels of BAC were determined in patients with liver diseases (N = 53), various cancers (N = 65) and other diseases (399). Serum levels of procollagen type III (PIIIP) amino propeptide, type IV collagen 7s domain (7s domain) and other parameters (TP, ALB, GOT, BPT, CHE, γ -GTP, ALP, LDH, CHE, TG, GLU) were also determined in those patients. In the whole patients, serum concns. of BAC showed a weak correlation with GOT, GPT, ALB and CHE but not with γ -GTP and ALP. There was no correlation between BAC and PIIIP or 7s domain. Although selum levels of BAC were elevated in both liver diseases and cancers, the increase in liver diseases was more marked. Markedly increased serum levels of BAC with low levels of CHE were found only in liver cirrhosis and liver cirrhosis plus hepatocellular carcinoma. Increased BAC may reflect caplillarization of the liver sinusoid or remodeling of the vascular basement membrane which is observed in the progression of liver fibrosis. Serum BAC is thought to be a promising new marker, different from PIIIP or 7s domain for diagnosing fibrosis state in the organs, particularly in the liver. vascular basement membrane collagen serum detn; disease basement ST membrane collagen detn ELISA; monoclonal antibody collagen detn disease ΙT Collagens, analysis RL: ANST (Analytical study) (determination of vascular basement membrane associated, of blood serum, by sandwich ELISA with monoclonal antibodies, in human diseases)

IT Disease

(serum vascular basement membrane associated collagen determination by

```
sandwich ELISA with monoclonal antibodies in, of
        humans)
IT
     Blood analysis
        (vascular basement membrane associated collagen determination in, by
        sandwich ELISA with monoclonal antibodies, in human
        diseases)
IT
     Liver, disease or disorder
        (fibrosis, serum vascular basement membrane associated collagen
        as marker for, in humans)
ΙT
     Antibodies
     RL: BIOL (Biological study)
        (monoclonal, for type IV collagen, in sandwich
        ELISA, serum vascular basememnt membrane associated
        collagen determination by, in human diseases)
L81 ANSWER 23 OF 29 HCAPLUS COPYRIGHT 2006 ACS on STN
AN
     1990:175179 HCAPLUS
DN
     112:175179
ED
     Entered STN: 12 May 1990
ΤI
     Determination of type IV collagen level in serum with monoclonal
     antibodies and its application to liver diseases
ΑU
     Kino, Jun; Yoshida, Tsuyoshi; Asamatsu, Chinatsu; Sato, Yoshihisa; Cho,
     Hiroko; Shiba, Kiyoko; Tomura, Shigeo; Hayashi, Toshihiko
     Shiseido Basic Res. Lab., Yokohama, 223, Japan
CS
SO
     Rinsho Kagaku (Nippon Rinsho Kagakkai) (1989), 18(4), 184-90
     CODEN: RIKAAN; ISSN: 0370-5633
DT
     Journal
LA
    English
     9-10 (Biochemical Methods)
CC
     Section cross-reference(s): 14, 15
AB
    A sandwich ELISA system was developed using 2
    monoclonal antibodies (JK-199 and JK-132) against type IV collagen
        Type IV collagen concentration in sera of healthy volunteers was 0
     .apprx. 65 ng/mL, but was elevated up to 480 ng/mL in sera of patients
     with liver cirrhosis. Type IV collagen concentration in sera of
     patients with liver disease showed no correlation with the serum concentration
of
    glutamate oxaloacetate transaminase, glutamate pyruvate transaminase,
    \gamma-glutamyl transpeptidase, and type III procollagen
    peptide (PIIIP). Either type IV collagen or PIIIP level was
     increased in sera from all the cases of hepatic carcinoma and/or liver
     cirrhosis compared to normal cases.
    type IV collagen detn patient ELISA; sandwich
    ELISA collagen detn blood patient; monoclonal antibody
    ELISA collagen patient; liver disease type IV
    collagen patient
ΙT
    Cirrhosis
    Diabetes mellitus
    Hepatitis
    Liver, neoplasm
        (type IV collagen determination in serum of, in human, by
        sandwich ELISA using monoclonal antibodies)
ΙT
    Blood analysis
        (type IV collagen determination in, by sandwich
        ELISA using monoclonal antibodies in patients with liver
        diseases or diabetes mellitus)
ΙŢ
    Antibodies
    RL: ANST (Analytical study)
        (monoclonal, for type IV collagen, in sandwich
        ELISA, in patients with liver diseases or diabetes mellitus)
```

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TΤ
     Collagens, analysis
     RL: ANT (Analyte); ANST (Analytical study)
        (type IV, determination of, by sandwich ELISA using
        monoclonal antibodies, in serum of patients with liver diseases or
        diabetes mellitus)
L81 ANSWER 24 OF 29 HCAPLUS COPYRIGHT 2006 ACS on STN
AN
     1990:174994 HCAPLUS
DN
     112:174994
ED
     Entered STN: 12 May 1990
ΤI
     A nonradioactive assay for type IV collagen degradation
ΑU
     Wilkinson, Mary J.; Cohen, Robert L.; Shuman, Marc A.
CS
     Cancer Res. Inst., Univ. California, San Francisco, CA, 94143, USA
     Analytical Biochemistry (1990), 185(2), 294-6
SO
     CODEN: ANBCA2; ISSN: 0003-2697
DT
     Journal
LA
     English
CC
     9-2 (Biochemical Methods)
     Section cross-reference(s): 7
AB
     A sensitive assay for type IV collagen degradation using an
     avidin-biotin sandwich technique is described. Biotinylated
     type IV collagen is allowed to bind to an avidin-coated
     microtiter plate. The solution to be assayed is incubated with the
     biotinylated collagen bound to the avidin plate.
     Collagen degraded by the solution is released into the supernatant
     and transferred to a second plate coated with avidin. By addition of
     biotinylated horseradish peroxidase to this second plate, the amount of
     collagen degraded is determined This assay requires only 0.5 µg of
     type IV collagen per microtiter plate and detects nanogram
     quantities of bacterial collagenase activity.
ST
     type IV collagen degrdn detn; collagenase detn avidin biotin
ΙT
     Bacteria
        (collagenase of, determination of, avidin-biotin sandwich technique
        for)
TT
     Avidins
     RL: ANST (Analytical study)
        (in type IV collagen degradation by sandwich technique)
ፐጥ
     Collagens, biological studies
     RL: PRP (Properties)
        (type IV, degradation of, avidin-biotin sandwich method for determination
        of)
TT
     9001-12-1, Collagenase
     RL: ANT (Analyte); ANST (Analytical study)
        (determination of, of bacteria, avidin-biotin sandwich technique for)
IT
     58-85-5, Biotin
     RL: ANST (Analytical study)
        (in type IV collagen degradation by sandwich technique)
L81
    ANSWER 25 OF 29 HCAPLUS COPYRIGHT 2006 ACS on STN
AN
     1990:95099 HCAPLUS
DN
     112:95099
ED
     Entered STN: 18 Mar 1990
     Human type IV collagen determination by sandwich EIA
ΤI
ΙN
     Obata, Kenichi; Iwata, Kazushi; Oshima, Akira; Inoue, Kyoichi
PΑ
     Fuji Chemicals Industrial Co., Ltd., Japan
     PCT Int. Appl., 59 pp.
SO
     CODEN: PIXXD2
DT
     Patent
LA
     Japanese
IC
     ICM G01N0033-53
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ICS G01N0033-577
CC
    9-10 (Biochemical Methods)
    Section cross-reference(s): 15
FAN.CNT 1
    PATENT NO.
                                    APPLICATION NO.
                     KIND DATE
                                                            DATE
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                      A1
ΡI
    WO 8907761
                             19890824 WO 1989-JP161
                                                            19890217 <--
        W: US
        RW: DE, FR, GB, IT
    JP 02001553
                A2
                            19900105 JP 1989-36111
                                                             19890217 <--
    JP 06077017
                      B4 19940928
    EP 401370
                      A1 19901212
                                        EP 1989-902540
                                                             19890217 <--
    EP 401370
                       В1
                            19950524
        R: DE, FR, GB, IT
JP 07072148 A2 19950317
PRAI JP 1988-35099 A 19880219
                                      JP 1993-252053
                                                        19930902 <--
                      A 19880219 <--
    WO 1989-JP161
                      W
                            19890217 <--
CLASS
            CLASS PATENT FAMILY CLASSIFICATION CODES
 PATENT NO.
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               ____
WO 8907761
              ICM G01N0033-53
               ICS G01N0033-577
               IPCI G01N0033-53 [ICM, 4]; G01N0033-577 [ICS, 4]
               IPCR C07K0016-18 [I,A]; C07K0016-18 [I,C]
 JP 02001553
               IPCI G01N0033-53 [ICM,5]; C12P0021-08 [ICS,5]; G01N0033-577
                     [ICS, 5]; A61K0039-395 [ICA, 5]
EP 401370
               IPCI G01N0033-53 [ICM, 5]; G01N0033-577 [ICS, 5]
               IPCR C07K0016-18 [I,A]; C07K0016-18 [I,C]
               ECLA
                      C07K016/18
 JP 07072148
               IPCI
                      G01N0033-53 [ICM, 6]; G01N0033-543 [ICS, 6]; G01N0033-577
                      [ICS, 6]
AΒ
    Sandwich EIA of the central triple helix moiety of human type IV
    collagen or that of human type IV collagen 7-S domain
    uses a monoclonal antibody capable of crossreacting with a specific moiety
    of the human type IV collagen. Thus, serum from chronic,
    inactive hepatitis patients was placed in a sensitized plate and incubated
    with peroxidase-labeled monoclonal antibody Fab' fragment from clone number
    1D3. Enzyme activity was measured for collagen determination Preparation of
    the monoclonal antibody is described.
ST
    type IV collagen sandwich EIA; serum type IV
    collagen ELISA hepatitis; monoclonal antibody type IV
    collagen immunoassay
TΤ
    Cirrhosis
    Liver, neoplasm
    Stomach, neoplasm
       (diagnosis of, type IV collagen determination in serum by
       sandwich EIA for)
ΙT
    Blood analysis
       (human type IV collagen determination in, by sandwich EIA,
       monoclonal antibody for)
IT
    Diagnosis
       (of liver cancer and other diseases, type IV collagen determination
       in serum by sandwich EIA for)
ΙT
    Hepatitis
       (chronic active, diagnosis of, type IV collagen determination in
       serum by sandwich EIA for)
IT
    Hepatitis
       (chronic persisting, diagnosis of, type IV collagen determination in
       serum by sandwich EIA for)
ΙT
    Antibodies
```

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RL: ANST (Analytical study)
        (monoclonal, to human type IV collagen, for sandwich
        EIA)
IT
     Collagens, analysis
     RL: ANST (Analytical study)
        (type IV, determination of human, in serum, by sandwich EIA,
        monoclonal antibody for)
L81 ANSWER 26 OF 29 HCAPLUS COPYRIGHT 2006 ACS on STN
     1989:474234 HCAPLUS
AN
     111:74234
DN
ΕD
     Entered STN: 03 Sep 1989
TТ
     One step sandwich enzyme immunoassay for human type IV
     collagen using monoclonal antibodies
ΑU
     Obata, Kenichi; Iwata, Kazushi; Ichida, Takafumi; Inoue, Kyoichi;
     Matsumoto, Eisaku; Muragaki, Yasuteru; Ooshima, Akira
CS
     Dep. Biotechnol., Fuji Chem. Ind., Ltd., Takaoka, 933, Japan
SO
     Clinica Chimica Acta (1989), 181(3), 293-303
     CODEN: CCATAR; ISSN: 0009-8981
DТ
     Journal
LA
     English
CC
     9-10 (Biochemical Methods)
     Section cross-reference(s): 14
AB
     Monoclonal antibodies were used in a one-step sandwich EIA for
     human serum immunoreactive type IV collagen. The one-step
     sandwich EIA using either polystyrene microspheres or microplates
     was characterized by carrying out two immunoreactions simultaneously, type
     IV collagen reacting with both a monoclonal antibody as a solid
     phase and a horseradish peroxidase-labeled monoclonal antibody (Fab')
     against human type IV collagen as a conjugate. The sensitivity
     with either polystyrene microspheres microplates was 0.22 ng per tube or
     0.04 ng per well for type IV collagen, and linearity was
     obtained at 0.22-40 ng/tube or 0.04-20 ng per well, resp.
     gave reproducible quant. anal. of immunoreactive type IV collagen
     levels in the sera of patients with hepatocellular carcinoma and patients
     with liver cirrhosis, which were higher than the levels in the sera of
     healthy subjects. Protein immunoblotting shows that the immunoreactive
     type IV collagen trapped in the title EIA system was not the 7-S
     and NC1 domains of type IV collagen.
ST
     serum type IV collagen detn; EIA type IV collagen;
     hepatocellular carcinoma serum collagen; liver cirrhosis serum
     collagen
IT
     Blood analysis
        (collagen type IV determination in, of human by EIA)
IT
     Cirrhosis
        (collagen type IV of human blood serum in)
IT
     Liver, neoplasm
        (hepatoma, collagen type IV of human blood serum in)
IT
     Antibodies
     RL: ANST (Analytical study)
        (monoclonal, to collagen type IV, for EIA)
IT
     Collagens, analysis
     RL: ANT (Analyte); ANST (Analytical study)
        (type IV, determination of, in human blood serum by EIA)
L81
    ANSWER 27 OF 29 HCAPLUS COPYRIGHT 2006 ACS on STN
ΑN
     1988:566876 HCAPLUS
DN
     109:166876
ED
     Entered STN: 12 Nov 1988
     EIA of human collagen peptides in blood for clinical diagnosis
ΤI
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IN
    Oshima, Akira; Iwata, Kazushi; Muragaki, Yasumitsu; Bai, Yasuo; Matsumoto,
     Eisaku; Miyamoto, Satoshi
PA
     Fuji Chemicals Industrial Co., Ltd., Japan
SO
     Jpn. Kokai Tokkyo Koho, 9 pp.
    CODEN: JKXXAF
DT
    Patent
LA
     Japanese
TC
     ICM G01N0033-543
     ICS G01N0033-577
CC
     9-10 (Biochemical Methods)
FAN.CNT 1
    PATENT NO.
                                     APPLICATION NO.
                       KIND
                               DATE
                                                               DATE
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                       ____
                              _____
                                          -----
PΤ
    JP 63063971
                       A2
                              19880322
                                          JP 1986-206862
                                                               19860904 <--
    JP 06038081
                       B4
                              19940518
                       A1
    CA 1287801
                              19910820
                                          CA 1987-530844
                                                               19870227 <--
                       Α
    US 5316914
                              19940531
                                          US 1992-831645
                                                               19920207 <--
PRAI JP 1986-206862
                       Α
                              19860904
                                       <--
    US 1987-22370
                       B1
                              19870305 <--
    US 1990-488440
                       В1
                             19900227
CLASS
 PATENT NO.
                CLASS PATENT FAMILY CLASSIFICATION CODES
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 JP 63063971
                ICM
                       G01N0033-543
                ICS
                       G01N0033-577
                IPCI
                       G01N0033-543 [ICM, 4]; G01N0033-577 [ICS, 4]
                IPCR
                       G01N0033-53 [N,A]; G01N0033-53 [N,C]; G01N0033-543
                       [I,A]; G01N0033-543 [I,C]; G01N0033-577 [I,A];
                       G01N0033-577 [I,C]
 CA 1287801
                IPCI
                       G01N0033-543 [ICM, 5]; G01N0033-577 [ICS, 5]
US 5316914
                IPCI
                       G01N0033-543 [ICM, 5]; G01N0033-576 [ICS, 5];
                       G01N0033-577 [ICS,5]
                IPCR
                       G01N0033-68 [I,A]; G01N0033-68 [I,C]
                NCL
                       435/007.940; 436/518.000; 436/548.000
AB
    A sandwich EIA for human type III, IV or VI collagen
    peptide determination uses immobilized monoclonal antibodies or polyclonal
    antibodies and enzyme-labeled monoclonal antibodies or polyclonal
    antibodies (at least one of the antibodies is a monoclonal antibody). A
    serum sample or standard was placed in a monoclonal antibody-sensitized
    microplate and incubated at room temperature for 1 h, followed by incubation
    with a rabbit polyclonal antibody and peroxidase-labeled goat antirabbit
    IgG (2nd antibody) at room temperature for 1 h and enzyme measurement for serum
    collagen peptide determination Blood collagen peptide levels in
    chronic active hepatitis and cirrhosis were markedly elevated.
ST
    collagen peptide detn serum EIA; liver disease diagnosis serum
    collagen
ΙT
    Blood analysis
        (collagen peptide determination in, by EIA)
ΙT
    Cirrhosis
       (diagnosis of, collagen peptide determination in blood serum by EIA
       for)
ΙT
    Peptides, analysis
    RL: ANST (Analytical study)
       (of collagen types III and IV and VI, determination of, in serum, by
       EIA)
ΙT
    Antibodies
    RL: ANST (Analytical study)
       (to collagen peptides III and IV and IV, for EIA)
ΙT
    Hepatitis
       (chronic active, diagnosis of, collagen peptide determination in
```

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blood serum by EIA for)
IT
     Immunochemical analysis
        (enzyme immunoassay, solid-phase, collagen peptide determination in
        blood serum by)
IT
     Antibodies
     RL: ANST (Analytical study)
        (monoclonal, to collagen peptides III and IV and IV, for EIA)
TT
     Collagens, analysis
     RL: ANST (Analytical study)
        (type III, peptide of, determination of, in serum, by EIA)
TΤ
     Collagens, analysis
     RL: ANST (Analytical study)
        (type IV, peptide of, determination of, in serum, by EIA)
TΤ
     Collagens, analysis
     RL: ANST (Analytical study)
        (type VI, peptide of, determination of, in serum, by EIA)
L81 ANSWER 28 OF 29 HCAPLUS COPYRIGHT 2006 ACS on STN
AN
     1982:611781 HCAPLUS
DN
     97:211781
ED
     Entered STN: 12 May 1984
ΤT
     Quantification of collagen types I and II in
     mouse limbs during differentiation in vitro and in vivo
ΑIJ
     Dusemund, B.; Barrach, H. J.
     Inst. Toxikol. Embryopharmakol., Freien Univ. Berlin, Berlin, 1000, Fed.
CS
     Rep. Ger.
SO
     Cult. Tech., Symp. Prenatal Dev., 5th (1981), 161-9. Editor(s):
     Neubert, Diether; Merker, Hans-Joachim. Publisher: de Gruyter, Berlin,
     Fed. Rep. Ger.
    CODEN: 48RRA8
DΤ
    Conference
LA
     English
CC
     9-2 (Biochemical Methods)
AΒ
     Collagens I and II were determined in developing mouse limbs in vitro
     and in vivo by a double-sandwich ELISA with
     peroxidase-conjugated antirabbit IgG. To determine the degree of
     collagen extraction from limb buds, the hydroxyproline contents of the
     total homogenates and the appropriate supernatants were compared, and the
     DNA concns. in the limbs were used as an indication of limb development.
     The method is sensitive, specific, and precise, and has the advantage of
     easy handling.
ST
     limb collagen detn differentiation; ELISA
     collagen limb differentiation; enzyme immunoassay collagen
     ; development limb collagen
IΤ
     Development, mammalian
        (collagens of limbs in)
IT
     Immunochemical analysis
        (enzyme-linked immunosorbent assay, for collagens)
IT
     Collagens, analysis
     RL: ANT (Analyte); ANST (Analytical study)
        (type I, determination of, in differentiating limbs by ELISA)
TT
     Collagens, analysis
     RL: ANT (Analyte); ANST (Analytical study)
        (type II, determination of, in differentiating limbs by
       ELISA)
IT
     51-35-4
     RL: ANT (Analyte); ANST (Analytical study)
        (determination of, in differentiating limbs, collagens determination in
        relation to)
```

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L81 ANSWER 29 OF 29 HCAPLUS COPYRIGHT 2006 ACS on STN
     1982:435489 HCAPLUS
AΝ
     97:35489
DN
ED
     Entered STN: 12 May 1984
TΙ
     Double-antibody enzyme-linked immunosorbent microassay for quantification
     of collagen types I and II
ΑU
     Dusemund, Birgit; Barrach, Hans Juergen
     Inst. Toxikol. Embryonalpharmakol., Freie Univ. Berlin, Berlin, D-1000,
CS
     Fed. Rep. Ger.
SO
     Journal of Immunological Methods (1982), 50(3), 255-68
     CODEN: JIMMBG; ISSN: 0022-1759
DT
     Journal
LA
     English
     9-2 (Biochemical Methods)
CC
AΒ
     Collagen types I and II were quantitated by
     sandwich enzyme-linked immunosorbent assay (ELISA) and
     double-sandwich ELISA. Specific anticollagen
     antibodies were linked to polystyrene microplates, and the
     collagen to be measured bound to the coating antibodies.
     Collagen type-specific 2nd antibodies reacted with the immobilized
     antigen.
              The 2nd antibodies were either labeled with peroxidase or were
     detected by using anti-IgG antibodies conjugated with peroxidase. Bound
     peroxidase was estimated by the color reaction produced with the substrate
     5-aminosalicylic acid. Optimization of the test procedure was achieved by
     varying the conditions for coating, antigen, and 2nd-antibody incubations.
     The detection limit with both methods was 0.033 \mug/mL, and the
     intra-assay relative standard deviations ranged 2.87-3.26 and 2.61-3.45% for
     the sandwich and double-sandwich ELISA,
     resp., for collagen type I. Similar results for precision,
     sensitivity, and specificity were obtained with both sandwich
     and double-sandwich ELISA. Both methods were more
     sensitive than inhibition ELISA and hydroxyproline determination
ST
     skin collagen detn; chondrosarcoma collagen detn;
     enzyme linked immunosorbent assay collagen
IT
     Skin, composition
        (collagen determination in, by double-antibody enzyme-linked
        immunosorbent assay)
ΤТ
     Sarcoma
        (chondro-, collagen determination in, by double-antibody
        enzyme-linked immunosorbent assay)
TΤ
     Immunochemical analysis
        (enzyme-linked immunosorbent assay, for collagen types)
TT
     Collagens, analysis
     RL: ANT (Analyte); ANST (Analytical study)
        (type I, determination of, by double-antibody enzyme-linked immunosorbent
        assay)
IT
     Collagens, analysis
     RL: ANT (Analyte); ANST (Analytical study)
        (type II, determination of, by double-antibody
        enzyme-linked immunosorbent assay)
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L1
                 E E15+ALL
L2
           67536 S E15+NT
         127133 S ?COLLAGEN?
L3
L4
          81362 S L1-L3 AND PY<=1997
L5
             157 S L4 AND SANDWICH
                 E SANDWICH/CT
                 E ELISA/CT
                 E E3+ALL
                 E E2+ALL
              30 S E67 AND L5
L6
L7
              32 S ELISA AND L5
              43 S L6, L7
L8
              1 S L8 AND 91011314/AN
L9
L10
             114 S L5 NOT L8
                 SEL AN 5 59 77 79 93 L10
               5 S E1-E5 AND L10
L11
L12
               6 S L9, L11
L13
              18 S L4 AND SANDWICH? NOT L5-L12
L14
               6 S L12 AND L1-L13
                 E QVIST/AU
L15
              51 S E29-E31
                 E ROSENQUIST/AU
L16
              49 S E14, E15, E18
                 E CHRISTGAU/AU
L17
              55 S E5, E6
              47 S L1-L3 AND L15-L17
L18
L19
              1 S L18 AND SANDWICH?
L20
             16 S L18 AND ELISA
                 E ELISA+ALL/CT
             19 S L18 AND E2+NT
L21
L22
             22 S L19-L21
L23
               7 S L22 AND PY<=1997
                 E CROSSLAP
L24
            193 S CROSSLAP?
L25
             36 S L24 AND PY<=1997
L26
              13 S L14, L23
              4 S L25 AND L26
L27
L28
             32 S L25 NOT L27
L29
             13 S L26, L27 AND L1-L28
     FILE 'MEDLINE' ENTERED AT 15:01:50 ON 11 MAY 2006
     FILE 'WPIX' ENTERED AT 15:02:19 ON 11 MAY 2006
L30
          16201 S ?COLLAGEN?
                 E COLLAGEN/CN
L31
               9 S E3-E5, E7-E12, E14-E18, E20-E24
                 SEL SDCN
                 EDIT E1-E9 /SDCN /DCN
L32
           2601 S E1-E9
                 SEL L31 DCSE
                 EDIT E10-E18 /DCSE /DCRE
L33
           2328 S E10-E18
L34
          16348 S L30, L32, L33
                 E A61K031-39/IC, ICM, ICS
                 E A61K038-39/IC, ICM, ICS
L35
            573 S E3-E10
                 E A61K038-39/ICA, ICI
L36
             16 S E3, E4
                 E A61K038:39/ICI
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L37
               6 S E3
L38
             572 S A61K038-39/IPC
                 E C07K014-78/IC, ICM, ICS
L39
             745 S E3-E5
                 E C07K014-78/ICA, ICI
              41 S E3, E4
1.40
                 E C07K014:78/ICI
               3 S E3
L41
                 E C07K014-78/IPC
             783 S C07K014-78/IPC
T.42
          16925 S L34-L42
L43
             87 S L43 AND SANDWICH?
L44
             189 S L43 AND ELISA
L45
L46
            111 S L43 AND ENZYM? (S) LINK? (S) IMMUNOSOR? (S) ASSAY?
L47
              13 S L44 AND L45, L46
                 SEL AN 8 12 L47
              2 S E1-E2 AND L47
L48
L49
              74 S L44 NOT L47
                 SEL AN 53 63
L50
               2 S E3-E4 AND L49
L51
               4 S L48, L50 AND L30-L50
     FILE 'WPIX' ENTERED AT 15:21:25 ON 11 MAY 2006
     FILE 'BIOSIS' ENTERED AT 15:21:36 ON 11 MAY 2006
                 E ROSENQUIST/AU
              46 S E13-E17
L52
                 E QVIST P/AU
              73 S E3-E5
L53
                 E CHRISTGAU S/AU
              74 S E3-E5
L54
L55
              68 S L52-L54 AND ?COLLAGEN?
L56
              8 S L55 AND PY<=1997
     FILE 'HCAPLUS' ENTERED AT 15:22:46 ON 11 MAY 2006
                 E COLLAGEN/CT
L57
            2136 S E21-E29
                 E E3+ALL
L58
           5398 S E1
                 E E2+ALL
          57099 S E3
L59
L60
            493 S E59-E62
L61
          62792 S L57-L60
L62
          94814 S COLLAGEN OR ?COLLAGEN OR ?COLLAGENS
L63
          56645 S L61, L62 AND (PY<=1997 OR PRY<=1997 OR AY<=1997)
L64
            121 S L63 AND SANDWICH?
L65
              45 S L64 AND (BIOCHEM?(L)METHOD?)/SC,SX
              8 S L65 AND TYPE()(II OR 2)
L66
L67
              8 S L65 AND TYPE(S) II
L68
              8 S L66, L67
L69
              37 S L65 NOT L68
                 E ELISA/CT
                 E E4+ALL
L70
          13817 S E2
L71
             50 S L63 AND L70
L72
            532 S L63 AND ELISA
L73
             32 S L64 AND L71, L72
L74
             27 S L73 NOT L68
L75
             56 S L69, L74
                 SEL AN L75 3 4 25 36 37 46-50 53
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FILE 'HCAPLUS' ENTERED AT 15:31:54 ON 11 MAY 2006

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